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=> file biosis caba caplus embase japio lifesci medline scisearch
=> e Norris steven j/au
E1
            8
                  NORRIS STEVEN/AU
E2
            3
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E3
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E4
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E5
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                  NORRIS STEVEN JAMES/AU
Ε6
            4
                  NORRIS STEVEN M/AU
Ε7
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Ε8
            1
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E9
            1
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E10
            3
                  NORRIS STEVEN R/AU
E11
            1
                  NORRIS STEVEN RANDOLPH/AU
E12
            3
                  NORRIS STUART/AU
=> s e3-e5 and borreli? and (VMP? or vls)
            47 ("NORRIS STEVEN J"/AU OR "NORRIS STEVEN J DR"/AU OR "NORRIS STEV
               EN JAMES"/AU) AND BORRELI? AND (VMP? OR VLS)
=> dup rem 11
PROCESSING COMPLETED FOR L1
             21 DUP REM L1 (26 DUPLICATES REMOVED)
=> d bib ab kwic 1-
YOU HAVE REQUESTED DATA FROM 21 ANSWERS - CONTINUE? Y/(N):y
    ANSWER 1 OF 21 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN
L2.
     DUPLICATE 1
ΑN
     2009:232172 BIOSIS <<LOGINID::20090609>>
DN
    PREV200900232172
ΤI
     Detailed Analysis of Sequence Changes Occurring during vlsE Antigenic
     Variation in the Mouse Model of ***Borrelia*** burgdorferi Infection.
ΑU
    Coutte, Loic [Reprint Author]; Botkin, Douglas J.; Gao, Lihui;
       ***Norris, Steven J.***
     Inst Biol Lille, Lille, France
CS
     Steven.J.Norris@uth.tmc.edu
SO
    PLoS Pathogens, (FEB 2009) Vol. 5, No. 2, pp. Article No.: e1000293.
    http://www.plospathogens.org.
     ISSN: 1553-7366. E-ISSN: 1553-7374.
     Article
DT
    English
LA
ED
     Entered STN: 1 Apr 2009
     Last Updated on STN: 1 Apr 2009
                  ***Borrelia*** can infect humans and animals for months
AB
     Lyme disease
     to years, despite the presence of an active host immune response. The
       ***vls***
                 antigenic variation system, which expresses the
     surface-exposed lipoprotein VlsE, plays a major role in B. burgdorferi
                                                           silent cassettes
     immune evasion. Gene conversion between
                                               ***vls***
     and the vlsE expression site occurs at high frequency during mammalian
     infection, resulting in sequence variation in the VlsE product. In this
     study, we examined vlsE sequence variation in B. burgdorferi B31 during
     mouse infection by analyzing 1,399 clones isolated from bladder, heart,
     joint, ear, and skin tissues of mice infected for 4 to 365 days. The
     median number of codon changes increased progressively in C3H/HeN mice
     from 4 to 28 days post infection, and no clones retained the parental vlsE
     sequence at 28 days. In contrast, the decrease in the number of clones
     with the parental vlsE sequence and the increase in the number of sequence
     changes occurred more gradually in severe combined immunodeficiency (SCID)
     mice. Clones containing a stop codon were isolated, indicating that
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continuous expression of full-length VlsE is not required for survival in vivo; also, these clones continued to undergo vlsE recombination. Analysis of clones with apparent single recombination events indicated that recombinations into vlsE are nonselective with regard to the silent cassette utilized, as well as the length and location of the recombination event. Sequence changes as small as one base pair were common. Fifteen percent of recovered vlsE variants contained "template-independent'' sequence changes, which clustered in the variable regions of vlsE. We hypothesize that the increased frequency and complexity of vlsE sequence changes observed in clones recovered from immunocompetent mice (as compared with SCID mice) is due to rapid clearance of relatively invariant clones by variable region-specific anti-VlsE antibody responses. Detailed Analysis of Sequence Changes Occurring during vlsE Antigenic Variation in the Mouse Model of ***Borrelia*** burgdorferi Infection. Coutte, Loic [Reprint Author]; Botkin, Douglas J.; Gao, Lihui;

- ΤI
- ΑIJ ***Norris, Steven J.***
- AΒ ***Borrelia*** can infect humans and animals for months Lyme disease to years, despite the presence of an active host immune response. ***vls*** antigenic variation system, which expresses the surface-exposed lipoprotein VlsE, plays a major role in B. burgdorferi immune evasion. Gene conversion between ***vls*** silent cassettes and the vlsE expression site occurs at high frequency during mammalian infection, resulting in sequence variation in the.
- ΙT system; bladder: excretory system; ear: sensory system; joint: skeletal system; skin tissue: integumentary system
- ΙT

Lyme disease: bacterial disease, ***Borrelia*** burgdorferi infection

Chemicals & Biochemicals IT

antibody

ORGN .

Mammals, Nonhuman Vertebrates, Nonhuman Mammals, Rodents, Vertebrates ORGN Classifier

> Spirochaetaceae 06112

Super Taxa

Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name

Borrelia burgdorferi (species): pathogen, strain-B31

Taxa Notes

Bacteria, Eubacteria, Microorganisms

- ANSWER 2 OF 21 L2 MEDLINE on STN
- ΑN 2006382551 MEDLINE <<LOGINID::20090609>>
- DN PubMed ID: 16796669
- ΤI Antigenic variation with a twist--the ***Borrelia*** story.
- ***Norris Steven J*** ΑU
- Department of Pathology. University of Texas Medical School at Houston, PO CS Box 20708, Houston, TX 77225-0708, USA.. Steven.J.Norris@uth.tmc.edu
- R01 AI37277 (United States NIAID NIH HHS) NC
- SO Molecular microbiology, (2006 Jun) Vol. 60, No. 6, pp. 1319-22. Journal code: 8712028. ISSN: 0950-382X.
- CY England: United Kingdom
- DTCommentary Journal; Article; (JOURNAL ARTICLE) (RESEARCH SUPPORT, N.I.H., EXTRAMURAL)
- English LA

FS Priority Journals EM200608 ΕD Entered STN: 27 Jun 2006 Last Updated on STN: 23 Aug 2006 Entered Medline: 22 Aug 2006 AΒ A common mechanism of immune evasion in pathogenic bacteria and protozoa is antigenic variation, in which genetic or epigenetic changes result in rapid, sequential shifts in a surface-exposed antigen. In this issue of Molecular Microbiology, Dai et al. provide the most complete description to date of the vlp/vsp antigenic variation system of the relapsing fever ***Borrelia*** hermsii. This elaborate, plasmid-encoded spirochaete, system involves an expression site that can acquire either variable large protein (vlp) or variable small protein (vsp) surface lipoprotein genes from 59 different archival copies. The archival vlp and vsp genes are arranged in clusters on at least five different plasmids. Gene conversion occurs through recombination events at upstream homology sequences (UHS) found in each gene copy, and at downstream homology sequences (DHS) found periodically among the vlp/vsp archival genes. Previous studies have shown that antigenic variation in relapsing fever ***Borrelia*** only permits the evasion of host antibody responses, but can also result in changes in neurotropism and other pathogenic properties. The ${\tt vlsE}$ antiqenic variation locus of Lyme disease spirochaetes, although similar in sequence to the relapsing fever vlp genes, has evolved a completely different antiqueic variation mechanism involving segmental recombination from a contiguous array of ***vls*** silent cassettes. These two systems thus appear to represent divergence from a common precursor followed by functional convergence to create two distinct antigenic variation processes. ΤI Antigenic variation with a twist--the ***Borrelia*** ΑU ***Norris Steven J*** AΒ . . et al. provide the most complete description to date of the vlp/vsp antigenic variation system of the relapsing fever spirochaete, ***Borrelia*** hermsii. This elaborate, plasmid-encoded system involves an expression site that can acquire either variable large protein (vlp) or variable small. . . homology sequences (DHS) found periodically among the vlp/vsp archival genes. Previous studies have shown that antigenic variation in relapsing fever ***Borrelia*** not only permits the evasion of host antibody responses, but can also result in changes in neurotropism and other pathogenic. . . relapsing fever vlp genes, has evolved a completely different antigenic variation mechanism involving segmental recombination from a contiguous array of ***vls*** silent cassettes. These two systems thus appear to represent divergence from a common precursor followed by functional convergence to create. . . *Antigenic Variation: GE, genetics CT *Antigens, Bacterial: GE, genetics *** Borrelia: GE, genetics*** ****Borrelia: IM, immunology*** L2 ANSWER 3 OF 21 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN 2007:87547 BIOSIS <<LOGINID::20090609>> ΑN PREV200700093298 DN ***VMP*** -like sequences of pathogenic TΙ ***Borrelia*** ΑU Anonymous; ***Norris, Steven J.*** [Inventor]; Zhang, Jing-Ren

[Inventor]; Hardham, John M. [Inventor]; Howell, Jerrilyn K. [Inventor];

Barbour, Alan G. [Inventor]; Weinstock, George M. [Inventor]

Houston, TX USA

CS

- ASSIGNEE: Board of Regents The University of Texas System
- PI US 07135176 20061114
- SO Official Gazette of the United States Patent and Trademark Office Patents, (NOV 14 2006)
 CODEN: OGUPE7. ISSN: 0098-1133.
- DT Patent
- LA English
- ED Entered STN: 31 Jan 2007 Last Updated on STN: 31 Jan 2007
- The present invention relates to DNA sequences encoding ***Vmp*** -like polypeptides of pathogenic ***Borrelia***, the use of the DNA sequences in recombinant vectors to express polypeptides, the encoded amino acid sequences, application of the DNA and amino acid sequences to the production of polypeptides as antigens for immunoprophylaxis, immunotherapy, and immunodiagnosis. Also disclosed are the use of the nucleic acid sequences as probes or primers for the detection of organisms causing Lyme disease, relapsing fever, or related disorders, and kits designed to facilitate methods of using the described polypeptides, DNA segments and antibodies.
- TI ***VMP*** -like sequences of pathogenic ***Borrelia*** .
- AU Anonymous; ***Norris, Steven J.*** [Inventor]; Zhang, Jing-Ren [Inventor]; Hardham, John M. [Inventor]; Howell, Jerrilyn K. [Inventor]; Barbour, Alan G. [Inventor]; Weinstock, George. . .
- AB The present invention relates to DNA sequences encoding ***Vmp*** -like polypeptides of pathogenic ***Borrelia*** , the use of the DNA sequences in recombinant vectors to express polypeptides, the encoded amino acid sequences, application of the. . .
- IT Major Concepts
 - Pharmacology; Clinical Immunology (Human Medicine, Medical Sciences); Infection
- IT Chemicals & Biochemicals
 - ***Borrelia*** ***VMP*** -like DNA sequences: diagnostic-drug, immunostimulant-drug, immunologic-drug
- L2 ANSWER 4 OF 21 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN DUPLICATE 2
- AN 2006:570967 BIOSIS <<LOGINID::20090609>>
- DN PREV200600576492
- TI Transcriptional regulation of the ***Borrelia*** burgdorferi antigenically variable VlsE surface protein.
- AU Bykowski, Tomasz; Babb, Kelly; von Lackum, Kate; Riley, Sean P.; ***Norris, Steven J.***; Stevenson, Brian [Reprint Author]
- CS Univ Kentucky, Coll Med, Dept Microbiol Mol Genet and Immunol, Albert B Chandler Med Ctr, MS 415, Lexington, KY 40536 USA brian.stevenson@uky.edu
- SO Journal of Bacteriology, (JUL 2006) Vol. 188, No. 13, pp. 4879-4889. CODEN: JOBAAY. ISSN: 0021-9193.
- DT Article
- LA English
- ED Entered STN: 1 Nov 2006 Last Updated on STN: 1 Nov 2006
- AB The Lyme disease agent ***Borrelia*** burgdorferi can persistently infect humans and other animals despite host active immune responses. This is facilitated, in part, by the vis locus, a complex system consisting of the vlsE expression site and an adjacent set of 11 to 15 silent ***vls*** cassettes. Segments of nonexpressed cassettes recombine with the vlsE region during infection of mammalian hosts,

resulting in combinatorial antigenic variation of the VISE outer surface protein. We now demonstrate that synthesis of VIsE is regulated during the natural mammal-tick infectious cycle, being activated in mammals but repressed during tick colonization. Examination of cultured B. burgdorferi cells indicated that the spirochete controls vIsE transcription levels in response to environmental cues. Analysis of PvIsE::gfp fusions in B. burgdorferi indicated that VIsE production is controlled at the level of transcriptional initiation, and regions of 5' DNA involved in the regulation were identified. Electrophoretic mobility shift assays detected qualitative and quantitative changes in patterns of protein-DNA complexes formed between the vIsE promoter and cytoplasmic proteins, suggesting the involvement of DNA-binding proteins in the regulation of vIsE, with at least one protein acting as a transcriptional activator.

- TI Transcriptional regulation of the ***Borrelia*** burgdorferi antigenically variable VlsE surface protein.
- AU Bykowski, Tomasz; Babb, Kelly; von Lackum, Kate; Riley, Sean P.; ***Norris, Steven J.***; Stevenson, Brian [Reprint Author]
- AB The Lyme disease agent ***Borrelia*** burgdorferi can persistently infect humans and other animals despite host active immune responses. This is facilitated, in part, by the. . . vis locus, a complex system consisting of the vlsE expression site and an adjacent set of 11 to 15 silent ***vls*** cassettes. Segments of nonexpressed cassettes recombine with the vlsE region during infection of mammalian hosts, resulting in combinatorial antigenic variation. . .

ORGN Notes

Animals, Chordates, Humans, Mammals, Primates, Vertebrates ORGN Classifier

Spirochaetaceae 06112

Super Taxa

Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name

Borrelia burgdorferi (species): pathogen

Taxa Notes

Bacteria, Eubacteria, Microorganisms

GEN ***Borrelia*** burgdorferi vlsE gene (Spirochaetaceae): expression

- L2 ANSWER 5 OF 21 CAPLUS COPYRIGHT 2009 ACS on STN DUPLICATE 3
- AN 2006:664489 CAPLUS <<LOGINID::20090609>>
- DN 145:183820
- TI Antiqenic variation with a twist: The ***Borrelia*** story
- AU ***Norris, Steven J.***
- CS Departments of Pathology & Laboratory Medicine and Microbiology & Molecular Genetics, University of Texas Medical School at Houston, Houston, TX, 77225-0708, USA
- SO Molecular Microbiology (2006), 60(6), 1319-1322 CODEN: MOMIEE; ISSN: 0950-382X
- PB Blackwell Publishing Ltd.
- DT Journal; General Review
- LA English
- AB A review. A common mechanism of immune evasion in pathogenic bacteria and protozoa is antigenic variation, in which genetic or epigenetic changes result in rapid, sequential shifts in a surface-exposed antigen. Dai et al. provide the most complete description to date of the vlp/vsp antigenic variation system of the relapsing fever spirochaete, ***Borrelia*** hermsii. This elaborate, plasmid-encoded system involves an expression

site that can acquire either variable large protein (vlp) or variable small protein (vsp) surface lipoprotein genes from 59 different archival copies. The archival vlp and vsp genes are arranged in clusters on at least five different plasmids. Gene conversion occurs through recombination events at upstream homol. sequences (UHS) found in each gene copy, and at downstream homol. sequences (DHS) found periodically among the vlp/vsp archival genes. Previous studies have shown that antigenic variation in relapsing fever ***Borrelia*** not only permits the evasion of host antibody responses, but can also result in changes in neurotropism and other pathogenic properties. The vlsE antigenic variation locus of Lyme disease spirochaetes, although similar in sequence to the relapsing fever vlp genes, has evolved a completely different antigenic variation mechanism involving segmental recombination from a contiquous array of ***vls*** silent cassettes. These two systems thus appear to represent divergence from a common precursor followed by functional convergence to create two distinct antigenic variation processes. RE.CNT 20 THERE ARE 20 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT Antigenic variation with a twist: The ***Borrelia*** ***Norris, Steven J.*** . . et al. provide the most complete description to date of the vlp/vsp antigenic variation system of the relapsing fever spirochaete, ***Borrelia*** hermsii. This elaborate, plasmid-encoded system

involves an expression site that can acquire either variable large protein (vlp) or variable small. . . homol. sequences (DHS) found periodically among the vlp/vsp archival genes. Previous studies have shown that antigenic variation in relapsing fever ***Borrelia*** not only permits the evasion of host antibody responses, but can also result in changes in neurotropism and other pathogenic. . . relapsing fever vlp genes, has evolved a completely different antigenic variation mechanism involving segmental recombination from a contiguous array of ***vls***

cassettes. These two systems thus appear to represent divergence from a common precursor followed by functional convergence to create.

review antigen ***Borrelia*** ST

Borrelia ΙT

(antigenic variation in ***Borrelia***)

ΤТ Antigens

ΤI

ΑU

AΒ

RL: BSU (Biological study, unclassified); BIOL (Biological study) (antigenic variation in ***Borrelia***)

ANSWER 6 OF 21 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN L2.

2006:134820 BIOSIS <<LOGINID::20090609>> AN

DN PREV200600145254

ТΤ ***Vmp*** -like sequences of pathogenic ***Borrelia***

Norris, Steven J. [Inventor]; Zhang, Jing-Ren [Inventor]; ΑU Hardham, John M. [Inventor]; Howell, Jerrilyn K. [Inventor]; Barbour, Alan G. [Inventor]; Weinstock, George M. [Inventor]

CS Houston, TX USA

ASSIGNEE: Board of Regents, The University of Texas System

US 06878816 20050412 PΙ

SO Official Gazette of the United States Patent and Trademark Office Patents, (APR 12 2005)

CODEN: OGUPE7. ISSN: 0098-1133.

DT Patent

English LA

```
Entered STN: 22 Feb 2006
ED
     Last Updated on STN: 22 Feb 2006
     The present invention relates to DNA sequences encoding ***Vmp*** -like
AΒ
     polypeptides of pathogenic ***Borrelia*** , the use of the DNA
     sequences in recombinant vectors to express polypeptides, the encoded
     amino acid sequences, application of the DNA and amino acid sequences to
     the production of polypeptides as antigens for immunoprophylaxis,
     immunotherapy, and immunodiagnosis. Also disclosed are the use of the
     nucleic acid sequences as probes or primers for the detection of organisms
     causing Lyme disease, relapsing fever, or related disorders, and kits
     designed to facilitate methods of using the described polypeptides, DNA
     segments and antibodies.
ΤI
       ***Vmp*** -like sequences of pathogenic
                                                ***Borrelia***
ΑU
       ***Norris, Steven J.***
                                [Inventor]; Zhang, Jing-Ren [Inventor];
     Hardham, John M. [Inventor]; Howell, Jerrilyn K. [Inventor]; Barbour, Alan
     G. [Inventor]; Weinstock, George. .
AΒ
     The present invention relates to DNA sequences encoding ***Vmp*** -like
     polypeptides of pathogenic
                                ***Borrelia*** , the use of the DNA
     sequences in recombinant vectors to express polypeptides, the encoded
     amino acid sequences, application of the. .
ΙT
        Clinical Immunology (Human Medicine, Medical Sciences); Infection;
        Clinical Chemistry (Allied Medical Sciences); Molecular Genetics
        (Biochemistry and Molecular Biophysics)
     Diseases
ΤT
            ***Borrelia*** infection: bacterial disease, drug therapy,
       prevention and control
ΙT
     Chemicals & Biochemicals
                        ***Vmp*** -like sequences;
                                                      ***Borrelia***
        DNA sequences;
        polypeptide antigens: diagnostic-drug, immunostimulant-drug,
        immunologic-drug
ORGN Classifier
                        06112
        Spirochaetaceae
     Super Taxa
        Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms
     Organism Name
           ***Borrelia*** (genus): pathogen
     Taxa Notes
        Bacteria, Eubacteria, Microorganisms
L2
    ANSWER 7 OF 21 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN
ΑN
     2004:283015 BIOSIS <<LOGINID::20090609>>
DN
     PREV200400283530
ΤI
       ***VMP*** -like sequences of pathogenic
                                                ***borrelia***
ΑU
       ***Norris, Steven J.*** [Inventor, Reprint Author]; Zhang, Jing-Ren
     [Inventor]; Hardham, John M. [Inventor]; Howell, Jerrilyn K. [Inventor];
     Barbour, Alan G. [Inventor]; Weinstock, George M. [Inventor]
CS
    ASSIGNEE: Board of Regents, The University of Texas System
PΙ
     US 6740744 20040525
    Official Gazette of the United States Patent and Trademark Office Patents,
SO
     (May 25 2004) Vol. 1282, No. 4.
     http://www.uspto.gov/web/menu/patdata.html. e-file.
     ISSN: 0098-1133 (ISSN print).
DT
    Patent
LA
    English
ED
    Entered STN: 9 Jun 2004
     Last Updated on STN: 9 Jun 2004
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```
The present invention relates to DNA sequences encoding ***Vmp*** -like
AΒ
     polypeptides of pathogenic ***Borrelia*** , the use of the DNA
     sequences in recombinant vectors to express polypeptides, the encoded
     amino acid sequences, application of the DNA and amino acid sequences to
     the production of polypeptides as antigens for immunoprophylaxis,
     immunotherapy, and immunodiagnosis. Also disclosed are the use of the
     nucleic acid sequences as probes or primers for the detection of organisms
     causing Lyme disease, relapsing fever, or related disorders, and kits
     designed to facilitate methods of using the described polypeptides, DNA
     segments and antibodies.
      ***VMP*** -like sequences of pathogenic
ΤI
                                                ***borrelia***
      ***Norris, Steven J.*** [Inventor, Reprint Author]; Zhang, Jing-Ren
ΑU
     [Inventor]; Hardham, John M. [Inventor]; Howell, Jerrilyn K. [Inventor];
     Barbour, Alan G. [Inventor]; . . .
AΒ
    The present invention relates to DNA sequences encoding ***Vmp*** -like
     polypeptides of pathogenic ***Borrelia*** , the use of the DNA
     sequences in recombinant vectors to express polypeptides, the encoded
     amino acid sequences, application of the.
ΤТ
    Major Concepts
       Equipment Apparatus Devices and Instrumentation; Infection; Methods and
        Techniques; Molecular Genetics (Biochemistry and Molecular Biophysics)
ΙT
    Diseases
           ***Borrelia*** infection: bacterial disease
           ***Borrelia*** Infections (MeSH)
ΤT
    Diseases
       Lyme disease: bacterial disease, diagnosis
       Lyme Disease (MeSH)
     Diseases
ΙT
       relapsing fever: bacterial disease, diagnosis
        Relapsing Fever (MeSH)
ΙT
    Chemicals & Biochemicals
            ***Vmp*** -like polypeptides: encoding DNA sequences, encoding
amino
       acid sequences; antibodies
ΙT
    Methods & Equipment
           ***Borrelia***
                           infection assay method: bioassay techniques,
        laboratory techniques; immunodiagnosis: immunologic techniques,
       laboratory techniques; immunoprophylaxis: immunologic techniques,
       laboratory techniques; immunotherapy: clinical techniques,. . .
ORGN Classifier
       Spirochaetaceae 06112
     Super Taxa
        Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms
     Organism Name
            ***Borrelia*** (genus): pathogen
     Taxa Notes
       Bacteria, Eubacteria, Microorganisms
L2
    ANSWER 8 OF 21 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN
     2004:257493 BIOSIS <<LOGINID::20090609>>
ΑN
DN
    PREV200400257602
TΙ
      ***VMP*** -like sequences of pathogenic ***Borrelia***
      ***Norris, Steven J.*** [Inventor, Reprint Author]; Zhang, Jing-Ren
ΑU
     [Inventor]; Hardham, John M. [Inventor]; Howell, Jerrilyn K. [Inventor];
    Barbour, Alan G. [Inventor]; Weinstock, George M. [Inventor]
CS
    Delmar, NY, USA
    ASSIGNEE: Board of Regents, The University of Texas System
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US 6719983 20040413
PΙ
SO
    Official Gazette of the United States Patent and Trademark Office Patents,
     (Apr 13 2004) Vol. 1281, No. 2.
     http://www.uspto.gov/web/menu/patdata.html. e-file.
     ISSN: 0098-1133 (ISSN print).
DT
    Patent
    English
LA
ED
    Entered STN: 12 May 2004
    Last Updated on STN: 12 May 2004
     The present invention relates to DNA sequences encoding ***Vmp*** -like
AΒ
    polypeptides of pathogenic ***Borrelia*** , the use of the DNA
     sequences in recombinant vectors to express polypeptides, the encoded
     amino acid sequences, application of the DNA and amino acid sequences to
     the production of polypeptides as antigens for immunoprophylaxis,
     immunotherapy, and immunodiagnosis. Also disclosed are the use of the
     nucleic acid sequences as probes or primers for the detection of organisms
     causing Lyme disease, relapsing fever, or related disorders, and kits
     designed to facilitate methods of using the described polypeptides, DNA
     segments and antibodies.
      ***VMP*** -like sequences of pathogenic ***Borrelia***
ΤI
      ***Norris, Steven J.*** [Inventor, Reprint Author]; Zhang, Jing-Ren
ΑU
     [Inventor]; Hardham, John M. [Inventor]; Howell, Jerrilyn K. [Inventor];
     Barbour, Alan G. [Inventor]; . . .
    The present invention relates to DNA sequences encoding ***Vmp*** -like
AΒ
    polypeptides of pathogenic ***Borrelia*** , the use of the DNA
     sequences in recombinant vectors to express polypeptides, the encoded
     amino acid sequences, application of the. .
ΙT
    Major Concepts
       Medical Genetics (Allied Medical Sciences); Molecular Genetics
        (Biochemistry and Molecular Biophysics)
ΙT
    Chemicals & Biochemicals
                             ***VMP*** -like DNA sequences
           ***Borrelia***
L2
    ANSWER 9 OF 21 CAPLUS COPYRIGHT 2009 ACS on STN
    2004:565053 CAPLUS <<LOGINID::20090609>>
ΑN
DN
    141:118336
    Polynucleotide and polypeptide sequences for ***vls*** genes of
ΤI
    pathogenic ***Borrelia*** and their diagnostic and therapeutic uses
    against infection and Lyme disease
     ***Norris, Steven J.***
IN
    Board of Regents, University of Texas System, USA
PΑ
    PCT Int. Appl., 182 pp.
SO
    CODEN: PIXXD2
DT
    Patent
LA
    English
FAN.CNT 1
                   KIND DATE
                                        APPLICATION NO.
    PATENT NO.
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    WO 2004058181
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                               20040715 WO 2003-US41182
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        RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ,
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BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE,
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            TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
                               20040722 AU 2003-299872
    AU 2003299872
                         Α1
                                                                 20031222
                        Α2
                               20050914
                                         EP 2003-800145
    EP 1572714
                                                                 20031222
           AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
            IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK
     US 20060240035
                       A1
                            20061026
                                         US 2005-539956
                                                                 20050617
PRAI US 2002-435077P
                         Ρ
                               20021220
     WO 2003-US41182
                         W
                               20031222
AB
    The invention claims DNA sequences encoding variable major protein (
      ***VMP*** )-like polypeptides of pathogenic ***Borrelia*** , the use
     of the DNA sequences in recombinant vectors to express polypeptides, the
     encoded amino acid sequences, application of the DNA and amino acid
     sequences to the prodn. of polypeptides as antigens for immunoprophylaxis,
     immunotherapy, and immunodiagnosis. The invention also claims use of the
     nucleic acid sequences as probes or primers for the detection of organisms
     causing Lyme disease, relapsing fever, or related disorders, and kits
     designed to facilitate methods of using the described polypeptides, DNA
     segments, and antibodies. Examples of the invention show reactivity of
     human Lyme disease serum with recombinant ***Borrelia*** afzelii
      ***Vls*** (variable major protein-like sequence) protein ***VLS***
     -BA13 and with recombinant B. garinii ***Vls*** protein ***VLS***
     -BG10. Mouse anti- ***Borrelia*** burgdorferi serum also reacted in an
     enzyme immunoassay with the recombinant proteins ***VLS*** -BA13 and
      ***VLS*** -BG10. The examples also show gene organization of
***vls***
     silent cassette loci from B. afzelii strain ACAI and B. garinii strain
     Ip90, expression of gene vlsE, and cDNA sequences of vlsE variants cloned
     from strains that were passaged through mice.
             THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD
             ALL CITATIONS AVAILABLE IN THE RE FORMAT
    Polynucleotide and polypeptide sequences for ***vls***
ΤI
    pathogenic
                ***Borrelia*** and their diagnostic and therapeutic uses
     against infection and Lyme disease
     ***Norris, Steven J.***
ΙN
     The invention claims DNA sequences encoding variable major protein (
AΒ
      of the DNA sequences in recombinant vectors to express polypeptides, the
     encoded amino acid sequences, application of the. . . the described
    polypeptides, DNA segments, and antibodies. Examples of the invention
     show reactivity of human Lyme disease serum with recombinant
      ***Borrelia*** afzelii ***Vls*** (variable major protein-like
                      ***VLS*** -BA13 and with recombinant B. garinii
     sequence) protein
      ***Vls***
                protein
                          ***VLS*** -BG10. Mouse anti- ***Borrelia***
     burgdorferi serum also reacted in an enzyme immunoassay with the
     recombinant proteins ***VLS*** -BA13 and ***VLS*** -BG10. The
     examples also show gene organization of ***vls*** silent cassette loci
     from B. afzelii strain ACAI and B. garinii strain Ip90, expression of gene
     vlsE, and cDNA sequences. .
    DNA sequence ***Borrelia*** gene ***vls*** antigen;

***Borrelia*** gene ***vls*** diagnosis vaccine immunotherapy Lyme
ST
    disease infection
ΙT
     Infection
       (bacterial; polynucleotide and polypeptide sequences for ***vls***
       genes of pathogenic ***Borrelia*** and their diagnostic and
       therapeutic uses against infection and Lyme disease)
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ΙT
     Immunoassay
        (enzyme-linked immunosorbent assay; polynucleotide and polypeptide
        sequences for ***vls*** genes of pathogenic
                                                        ***Borrelia***
        their diagnostic and therapeutic uses against infection and Lyme
       disease)
ΤТ
    Recombination, genetic
        (gene conversion; polynucleotide and polypeptide sequences for
          ***vls*** genes of pathogenic ***Borrelia*** and their
diagnostic
        and therapeutic uses against infection and Lyme disease)
ΙT
     Diagnosis
        (immunodiagnosis; polynucleotide and polypeptide sequences for
          ***vls*** genes of pathogenic ***Borrelia*** and their
diagnostic
        and therapeutic uses against infection and Lyme disease)
ΤТ
    Animals
     Bos taurus
     Canis familiaris
     Cervidae
     Equus caballus
     Human
    Mus
        (infection; polynucleotide and polypeptide sequences for ***vls***
        genes of pathogenic ***Borrelia*** and their diagnostic and
        therapeutic uses against infection and Lyme disease)
ΤT
    Antibodies and Immunoglobulins
     RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); DGN
     (Diagnostic use); ANST (Analytical study); BIOL (Biological study); USES
     (Uses)
        (labeled; polynucleotide and polypeptide sequences for
                                                               ***vls***
                             ***Borrelia***
        genes of pathogenic
                                              and their diagnostic and
        therapeutic uses against infection and Lyme disease)
     Antibodies and Immunoglobulins
ΤT
     RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); DGN
     (Diagnostic use); ANST (Analytical study); BIOL (Biological study); USES
     (Uses)
        (monoclonal; polynucleotide and polypeptide sequences for
        genes of pathogenic ***Borrelia*** and their diagnostic and
        therapeutic uses against infection and Lyme disease)
ΙT
     Antigenic variation
     Blood analysis
         ***Borrelia***
                        afzelii
         ***Borrelia*** burgdorferi
                        garinii
         ***Borrelia***
     DNA sequences
     Genetic polymorphism
     Immunity
     Immunoassay
     Immunoblotting
     Immunoprecipitation
     Immunotherapy
     Lyme disease
     Molecular cloning
     Nucleic acid amplification (method)
     Plasmids
     Protein sequences
     Radioimmunoassay
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Test kits
     Urine analysis
     cDNA sequences
        (polynucleotide and polypeptide sequences for ***vls***
                   ***Borrelia*** and their diagnostic and therapeutic uses
       pathogenic
       against infection and Lyme disease)
ΙT
    Antigens
    RL: ANT (Analyte); BPN (Biosynthetic preparation); DGN (Diagnostic use);
     PRP (Properties); ANST (Analytical study); BIOL (Biological study); PREP
     (Preparation); USES (Uses)
        (polynucleotide and polypeptide sequences for ***vls***
       pathogenic ***Borrelia*** and their diagnostic and therapeutic uses
       against infection and Lyme disease)
ΙT
    Nucleic acids
    RNA
    RL: ANT (Analyte); BUU (Biological use, unclassified); DGN (Diagnostic
     use); ANST (Analytical study); BIOL (Biological study); USES (Uses)
        (polynucleotide and polypeptide sequences for ***vls***
                   ***Borrelia*** and their diagnostic and therapeutic uses
       pathogenic
       against infection and Lyme disease)
ΙT
    Antibodies and Immunoglobulins
    RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); DGN
     (Diagnostic use); ANST (Analytical study); BIOL (Biological study); USES
     (Uses)
        (polynucleotide and polypeptide sequences for ***vls***
       pathogenic ***Borrelia*** and their diagnostic and therapeutic uses
       against infection and Lyme disease)
ΙT
    Primers (nucleic acid)
     RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); DGN
     (Diagnostic use); PRP (Properties); ANST (Analytical study); BIOL
     (Biological study); USES (Uses)
        (polynucleotide and polypeptide sequences for ***vls***
                                                                   genes of
                    ***Borrelia*** and their diagnostic and therapeutic uses
       pathogenic
       against infection and Lyme disease)
ΙT
    Escherichia coli
        (recombinant host; polynucleotide and polypeptide sequences for
          ***vls*** genes of pathogenic ***Borrelia*** and their
diagnostic
       and therapeutic uses against infection and Lyme disease)
ΙT
     Fever and Hyperthermia
        (relapsing; polynucleotide and polypeptide sequences for ***vls***
       genes of pathogenic ***Borrelia*** and their diagnostic and
       therapeutic uses against infection and Lyme disease)
ΙT
    Repetitive DNA
     RL: BSU (Biological study, unclassified); PRP (Properties); BIOL
     (Biological study)
        ( ***vls*** silent cassettes; polynucleotide and polypeptide
       sequences for ***vls*** genes of pathogenic ***Borrelia***
                                                                          and
       their diagnostic and therapeutic uses against infection and Lyme
       disease)
IT
    Gene, microbial
    RL: ANT (Analyte); BUU (Biological use, unclassified); DGN (Diagnostic
     use); PRP (Properties); ANST (Analytical study); BIOL (Biological study);
    USES (Uses)
        ( ***vls*** ; polynucleotide and polypeptide sequences for
          ***vls*** genes of pathogenic ***Borrelia*** and their
diagnostic
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and therapeutic uses against infection and Lyme disease)
ΤТ
    Gene, microbial
     RL: ANT (Analyte); BUU (Biological use, unclassified); DGN (Diagnostic
     use); PRP (Properties); ANST (Analytical study); BIOL (Biological study);
     USES (Uses)
       (vlsE; polynucleotide and polypeptide sequences for ***vls*** genes
       of pathogenic ***Borrelia*** and their diagnostic and therapeutic
       uses against infection and Lyme disease)
ΙT
    721865-74-3 721865-75-4 721865-91-4 721865-92-5
    RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); DGN
     (Diagnostic use); PRP (Properties); ANST (Analytical study); BIOL
     (Biological study); USES (Uses)
       ( ***Borrelia*** afzelii strain ACAI gene vls13 primer;
       polynucleotide and polypeptide sequences for ***vls*** genes of
       pathogenic ***Borrelia*** and their diagnostic and therapeutic uses
       against infection and Lyme disease)
ΙT
    721865-72-1
    RL: ANT (Analyte); BUU (Biological use, unclassified); DGN (Diagnostic
     use); PRP (Properties); ANST (Analytical study); BIOL (Biological study);
     USES (Uses)
       ( ***Borrelia*** burgdorferi B31 vlsE and ***vls*** silent
       cassette flanking direct repeat; polynucleotide and polypeptide
       sequences for ***vls*** genes of pathogenic ***Borrelia***
       their diagnostic and therapeutic uses against infection and Lyme
       disease)
    721865-93-6
                  721865-94-7 721865-95-8
                                            721865-96-9
ΤТ
     RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); DGN
     (Diagnostic use); PRP (Properties); ANST (Analytical study); BIOL
     (Biological study); USES (Uses)
       ( ***Borrelia*** garinii strain Ip90 gene vls10 primer;
       polynucleotide and polypeptide sequences for ***vls*** genes of
       pathogenic ***Borrelia*** and their diagnostic and therapeutic uses
       against infection and Lyme disease)
ΙT
    721863-14-5
     RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); DGN
     (Diagnostic use); PRP (Properties); ANST (Analytical study); BIOL
     (Biological study); USES (Uses)
       ( ***Borrelia*** gene
                                 ***vls*** specific primer 4470;
       polynucleotide and polypeptide sequences for ***vls*** genes of
       pathogenic ***Borrelia***
                                   and their diagnostic and therapeutic uses
       against infection and Lyme disease)
    721863-15-6
ΤT
     RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); DGN
     (Diagnostic use); PRP (Properties); ANST (Analytical study); BIOL
     (Biological study); USES (Uses)
                          gene ***vls***
       ( ***Borrelia***
                                             specific primer 4471;
       polynucleotide and polypeptide sequences for ***vls*** genes of
       pathogenic ***Borrelia*** and their diagnostic and therapeutic uses
       against infection and Lyme disease)
ΤТ
    721863-03-2
    RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); DGN
     (Diagnostic use); PRP (Properties); ANST (Analytical study); BIOL
     (Biological study); USES (Uses)
       ( ***Borrelia*** gene ***vls*** specific primer 4540;
       polynucleotide and polypeptide sequences for ***vls*** genes of
       pathogenic ***Borrelia*** and their diagnostic and therapeutic uses
       against infection and Lyme disease)
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721863-11-2
TТ
    RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); DGN
     (Diagnostic use); PRP (Properties); ANST (Analytical study); BIOL
     (Biological study); USES (Uses)
                         gene ***vls***
       ( ***Borrelia***
                                           specific primer 4545;
       polynucleotide and polypeptide sequences for ***vls*** genes of
       pathogenic ***Borrelia*** and their diagnostic and therapeutic uses
       against infection and Lyme disease)
ΙT
    721863-10-1
    RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); DGN
     (Diagnostic use); PRP (Properties); ANST (Analytical study); BIOL
     (Biological study); USES (Uses)
       ( ***Borrelia*** gene ***vls*** specific primer 4548;
       polynucleotide and polypeptide sequences for ***vls*** genes of
       pathogenic ***Borrelia*** and their diagnostic and therapeutic uses
       against infection and Lyme disease)
ΙT
    721863-12-3
    RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); DGN
     (Diagnostic use); PRP (Properties); ANST (Analytical study); BIOL
     (Biological study); USES (Uses)
       ( ***Borrelia*** gene ***vls*** specific primer 4587;
       polynucleotide and polypeptide sequences for ***vls*** genes of
       pathogenic ***Borrelia*** and their diagnostic and therapeutic uses
       against infection and Lyme disease)
ΤТ
    721863-13-4
    RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); DGN
    (Diagnostic use); PRP (Properties); ANST (Analytical study); BIOL
     (Biological study); USES (Uses)
       ( ***Borrelia***
                                ***vls*** specific primer 4588;
                         gene
       polynucleotide and polypeptide sequences for ***vls*** genes of
                   ***Borrelia*** and their diagnostic and therapeutic uses
       pathogenic
       against infection and Lyme disease)
    721862-92-6P 721862-95-9P 721863-00-9P 721863-01-0P
                                                              721863-02-1P
ΤT
    721863-07-6P 721863-08-7P 721863-09-8P 721863-19-0P 721863-20-3P
    721863-33-8P 721863-34-9P 721863-35-0P 721863-36-1P 721863-37-2P
    721863-38-3P 721863-39-4P 721863-40-7P 721863-41-8P 721863-42-9P
    721863-43-0P 721863-45-2P
                                 721863-48-5P 721863-62-3P
                                                              721863-63-4P
    721863-64-5P 721863-65-6P
                                721863-66-7P 721863-67-8P
                                                             721863-68-9P
    721863-70-3P 721863-73-6P 721863-74-7P 721865-61-8P, Antigen
    (plasmid pBG-10-1 gene vls10) 721865-63-0P 721865-64-1P 721865-65-2P
    721865-66-3P 721865-67-4P 721865-68-5P 721865-71-0P, Antigen
    (plasmid pBA-13-1 gene vls13)
    RL: ANT (Analyte); BPN (Biosynthetic preparation); DGN (Diagnostic use);
    PRP (Properties); ANST (Analytical study); BIOL (Biological study); PREP
     (Preparation); USES (Uses)
       (amino acid sequence; polynucleotide and polypeptide sequences for
         ***vls*** genes of pathogenic ***Borrelia*** and their
diagnostic
       and therapeutic uses against infection and Lyme disease)
ΙT
    511612-64-9 511612-65-0 511612-66-1 511612-67-2 511612-68-3
    511612-69-4 511612-70-7 511612-71-8 511612-72-9
                                                         511612-73-0
    511612-74-1 511612-75-2 511612-76-3 511612-77-4
                                                          511612-78-5
    511612-79-6, Antigen ( ***Borrelia*** afzelii strain ACAI clone 2622
    gene vlsE C-terminal fragment) 511612-80-9, Antigen ( ***Borrelia***
    afzelii strain ACAI clone 2624a gene vlsE C-terminal fragment)
    511612-81-0, Antigen ( ***Borrelia*** afzelii strain ACAI clone 2624b
    gene vlsE C-terminal fragment) 511612-82-1, Antigen ( ***Borrelia***
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afzelii strain ACAI clone 2625 gene vlsE fragment) 511612-83-2
     511612-84-3 511612-85-4 511612-86-5 511612-87-6 511612-88-7
    511612-89-8 511612-90-1 511612-91-2 511612-92-3 511612-93-
511612-94-5 511612-95-6 511612-96-7, Antigen ( ***Borrelia***
                                                           511612-93-4
     garinii strain Ip90 clone 17 gene vlsE fragment)
                                                       511612-97-8, Antigen (
      ***Borrelia*** garinii strain Ip90 clone 20 gene vlsE fragment)
     511612-98-9, Antigen ( ***Borrelia*** garinii strain Ip90 clone 21 gene
     vlsE fragment) 511612-99-0, Antigen ( ***Borrelia*** garinii strain
     Ip90 clone 23 gene vlsE fragment)
     RL: BSU (Biological study, unclassified); PRP (Properties); BIOL
     (Biological study)
        (amino acid sequence; polynucleotide and polypeptide sequences for
          ***vls*** genes of pathogenic ***Borrelia*** and their
diagnostic
       and therapeutic uses against infection and Lyme disease)
    721862-91-5 721862-96-0 721862-97-1 721862-98-2
ΤT
                                                          721862-99-3
                              721863-06-5
     721863-04-3
                  721863-05-4
                                              721863-16-7
                                                            721863-21-4
                 721863-23-6 721863-24-7
     721863-22-5
                                             721863-25-8 721863-26-9
    721863-27-0 721863-28-1 721863-29-2 721863-30-5 721863-31-6
    721863 - 32 - 7 721863 - 44 - 1 721863 - 46 - 3 721863 - 47 - 4 721863 - 49 - 6
    721863-50-9 721863-51-0 721863-52-1 721863-53-2 721863-54-3
    721863-55-4 721863-56-5 721863-57-6 721863-58-7
                                                          721863-59-8
                  721863-61-2 721863-69-0 721863-71-4
                                                            721863-72-5
     721863-60-1
    721865-60-7, DNA (plasmid pBG-10-1 gene vls10) 721865-62-9, DNA (plasmid
    pBA-13-1 gene vls13) 721865-69-6 721865-70-9
    RL: ANT (Analyte); BUU (Biological use, unclassified); DGN (Diagnostic
     use); PRP (Properties); ANST (Analytical study); BIOL (Biological study);
    USES (Uses)
        (nucleotide sequence; polynucleotide and polypeptide sequences for
         ***vls*** genes of pathogenic ***Borrelia*** and their
diagnostic
       and therapeutic uses against infection and Lyme disease)
     503713-49-3 503713-50-6 503713-51-7 503713-52-8 503713-53-9
ΤT
     503713-54-0
                  503713-55-1 503713-56-2 503713-57-3
                                                            503713-58-4
    RL: BSU (Biological study, unclassified); PRP (Properties); BIOL
     (Biological study)
        (nucleotide sequence; polynucleotide and polypeptide sequences for
          ***vls*** genes of pathogenic ***Borrelia*** and their
diagnostic
       and therapeutic uses against infection and Lyme disease)
     58-85-5D, Biotin, conjugates
    RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); DGN
     (Diagnostic use); ANST (Analytical study); BIOL (Biological study); USES
        (polynucleotide and polypeptide sequences for ***vls***
                                                                   genes of
                   ***Borrelia***
                                    and their diagnostic and therapeutic uses
       pathogenic
       against infection and Lyme disease)
     145856-09-3, GenBank L04788 391840-97-4, GenBank U76405
ΤТ
     RL: BSU (Biological study, unclassified); PRP (Properties); BIOL
     (Biological study)
        (polynucleotide and polypeptide sequences for ***vls*** genes of
       pathogenic ***Borrelia*** and their diagnostic and therapeutic uses
       against infection and Lyme disease)
ΤТ
    721865-73-2
    RL: BUU (Biological use, unclassified); PRP (Properties); BIOL (Biological
     study); USES (Uses)
        (restriction endonuclease EcoRI-site linker; polynucleotide and
```

polypeptide sequences for ***vls*** genes of pathogenic ***Borrelia*** and their diagnostic and therapeutic uses against infection and Lyme disease) 721869-22-3 721869-24-5 ΙT 721869-20-1 RL: PRP (Properties) (unclaimed nucleotide sequence; polynucleotide and polypeptide sequences for ***vls*** genes of pathogenic ***Borrelia*** their diagnostic and therapeutic uses against infection and Lyme disease) ΙT 721869-21-2 721869-23-4 RL: PRP (Properties) (unclaimed protein sequence; polynucleotide and polypeptide sequences ***vls*** genes of pathogenic ***Borrelia*** and their diagnostic and therapeutic uses against infection and Lyme disease) ΤТ 116934-33-9 RL: PRP (Properties) (unclaimed sequence; polynucleotide and polypeptide sequences for ***vls*** genes of pathogenic ***Borrelia*** diagnostic and therapeutic uses against infection and Lyme disease) ANSWER 10 OF 21 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on L2 DUPLICATE 4 2005:50985 BIOSIS <<LOGINID::20090609>> ΑN PREV200500047406 DN ΤI Effects of vlsE complementation on the infectivity of ***Borrelia*** burgdorferi lacking the linear plasmid 1p28-1. ΑU Lawrenz, Matthew B.; Wooten, R. Mark; ***Norris, Steven J.*** [Reprint Author] CS Sch MedDept Pathol and Lab Med, Univ Texas, POB 20708, Houston, TX, 77225, steven.j.norris@uth.tmc.edu Infection and Immunity, (November 2004) Vol. 72, No. 11, pp. 6577-6585. SO ISSN: 0019-9567 (ISSN print). DT Article LA English Entered STN: 26 Jan 2005 ED Last Updated on STN: 26 Jan 2005 The loss of linear plasmid lp28-1, which contains the ***vls*** AB antigenic variation locus, is associated with reduced infectivity of ***Borrelia*** burgdorfieri in immunocompetent mice. The recombinant shuttle vector pBBE22, which includes the virulence determinant BBE22 from 1p25 and restores infectivity to readily transformable B. burgdorferi lacking 1p25 and 1p56, was used to determine the effect of trans expression of vlsE on virulence. Spirochetes lacking lp28-1 were complemented with the plasmid pBBE22:vlsE, containing both BBE22 and vlsE. VlsE protein produced by this construct was expressed and surface accessible in in vitro-cultured B. burgdorferi, as determined by surface proteolysis and immunoblot analysis. Clones lacking 1p25 but containing 1p28-1 and either pBBE22 or pBBE22:vlsE were reisolated consistently from immunocompetent mice 8 weeks after infection. In contrast, a clone

lacking both 1p25 and 1p28-1 and complemented with pBBE22:vlsE was isolated from only a single tissue of one of six C3H/HeN mice 8 weeks postinfection. These results indicate that either an intact v/s antigenic variation locus or another determinant on 1p28-1 is required to restore complete infectivity. In addition, an isogenic clone that retained 1p28-1

was complemented with the v/sE shuttle plasmid and was examined for vlsE sequence variation and infectivity. Sequence variation was not observed for the shuttle plasmid, indicating that the cis arrangement of v/sE and the ***vls*** silent cassettes in 1p28-1 facilitate vlsE gene conversion. Lack of vlsE sequence variation on the shuttle plasmid thus did not result in clearance of the trans-complemented strain in immunocompetent mice under the conditions tested.

- TI Effects of vlsE complementation on the infectivity of ***Borrelia*** burgdorferi lacking the linear plasmid lp28-1.
- AU Lawrenz, Matthew B.; Wooten, R. Mark; ***Norris, Steven J.*** [Reprint Author]
- AB The loss of linear plasmid lp28-1, which contains the ***vls*** antigenic variation locus, is associated with reduced infectivity of ***Borrelia*** burgdorfieri in immunocompetent mice. The recombinant shuttle vector pBBE22, which includes the virulence determinant BBE22 from lp25 and restores infectivity. . . and infectivity. Sequence variation was not observed for the shuttle plasmid, indicating that the cis arrangement of v/sE and the ***vls*** silent cassettes in lp28-1 facilitate vlsE gene conversion. Lack of vlsE sequence variation on the shuttle plasmid thus did not. . .

ORGN . . .

Mammals, Nonhuman Vertebrates, Nonhuman Mammals, Rodents, Vertebrates ORGN Classifier

Spirochaetaceae 06112

Super Taxa

Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name

Borrelia burgdorferi (species): pathogen

Taxa Notes

Bacteria, Eubacteria, Microorganisms

- L2 ANSWER 11 OF 21 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN DUPLICATE 5
- AN 2003:153698 BIOSIS <<LOGINID::20090609>>
- DN PREV200300153698
- TI Characterization of the ***vls*** antigenic variation loci of the Lyme disease spirochaetes ***Borrelia*** garinii Ip90 and ***Borrelia*** afzelii ACAI.
- AU Wang, Dachun; Botkin, Douglas J.; ***Norris, Steven J.*** [Reprint Author]
- CS Department of Pathology and Laboratory Medicine, Medical School at Houston, University of Texas, PO Box 20708, Houston, TX, 77225-0708, USA Steven.J.Norris@uth.tmc.edu
- SO Molecular Microbiology, (March 2003) Vol. 47, No. 5, pp. 1407-1417. print. ISSN: 0950-382X (ISSN print).
- DT Article
- LA English
- ED Entered STN: 26 Mar 2003 Last Updated on STN: 26 Mar 2003
- AB The ***vls*** locus of ***Borrelia*** burgdorferi B31 consists of 15 silent cassettes and one expression site (vlsE), and the presence of the encoding plasmid Ip28-1 correlates with high infectivity.

 Recombination between the expression cassette and silent cassettes occurs in vivo, and this process may enable B. burgdorferi to evade the immune response. To determine the characteristics of the ***vls*** loci in other ***Borrelia*** strains, we have cloned and characterized the ***vls*** silent cassette loci of ***Borrelia*** garinii Ip90 and

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***Borrelia*** afzelii ACAI, consisting of 11 ***vls*** silent
    cassettes and 14 ***vls*** silent cassettes respectively. The silent
    cassettes share 90-97% nucleotide sequence identity with one another
    within the Ip90 ***vls*** locus and 84-91% within the ACAI
                                                                   ***vls***
    locus. In both organisms, the silent cassettes resemble the B31
      ***Vls*** sequences in overall amino acid similarity (50-65%) and in
    presence of six variable regions interspersed between six relatively
    invariant regions. The vise expression sites of these two strains have
    not been isolated, but transcripts of vise were detected by reverse
    transcriptase-polymerase chain reaction for both Ip90 and ACAI. In
    addition, the occurrence of sequence variation within the vlsE cassette
    region of these transcripts was verified. This study indicates that the
                loci present in B. garinii Ip90 and B. afzelii ACAI have
    characteristics similar to those found in B. burgdorferi B31.
    Characterization of the ***vls*** antigenic variation loci of the Lyme
    disease spirochaetes ***Borrelia*** garinii Ip90 and ***Borrelia***
    afzelii ACAI.
    Wang, Dachun; Botkin, Douglas J.; ***Norris, Steven J.*** [Reprint
         ***vls*** locus of ***Borrelia*** burgdorferi B31 consists of
    The
    15 silent cassettes and one expression site (vlsE), and the presence of
    the encoding plasmid Ip28-1. . . in vivo, and this process may enable
    B. burgdorferi to evade the immune response. To determine the
    characteristics of the ***vls*** loci in other ***Borrelia***
    strains, we have cloned and characterized the ***vls*** silent
    cassette loci of ***Borrelia*** garinii Ip90 and
                                                        ***Borrelia***
    afzelii ACAI, consisting of 11 ***vls*** silent cassettes and 14
      ***vls*** silent cassettes respectively. The silent cassettes share
    90-97% nucleotide sequence identity with one another within the Ip90
      ***vls*** locus and 84-91% within the ACAI ***vls*** locus. In
both
    organisms, the silent cassettes resemble the B31 ***Vls***
    in overall amino acid similarity (50-65\%) and in the presence of six
    variable regions interspersed between six relatively invariant. . . the
    occurrence of sequence variation within the vlsE cassette region of these
    transcripts was verified. This study indicates that the ***vls***
    loci present in B. garinii Ip90 and B. afzelii ACAI have characteristics
    similar to those found in B. burgdorferi B31.
ORGN Classifier
       Spirochaetaceae 06112
    Super Taxa
       Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms
    Organism Name
           ***Borrelia*** afzelii (species): parasite, strain-ACAI
           ***Borrelia*** burgdorferi (species): parasite, B31
           ***Borrelia*** garinii (species): parasite, strain-Ip90
    Taxa Notes
       Bacteria, Eubacteria, Microorganisms
      ***vls***
                 gene: antigenic variation loci
    ANSWER 12 OF 21 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on
    2002:523404 BIOSIS <<LOGINID::20090609>>
    PREV200200523404
      ***VMP*** -like sequences of pathogenic ***borrelia*** .
      ***Norris, Steven J.*** [Inventor, Reprint author]; Zhang, Jing-Ren
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[Inventor]; Hardham, John M. [Inventor]; Howell, Jerrilyn K. [Inventor]; Barbour, Alan G. [Inventor]; Weinstock, George M. [Inventor] Houston, TX, USA ASSIGNEE: Board of Regents, The University of Texas System US 6437116 20020820 Official Gazette of the United States Patent and Trademark Office Patents, (Aug. 20, 2002) Vol. 1261, No. 3. http://www.uspto.gov/web/menu/patdata.html. e-file. CODEN: OGUPE7. ISSN: 0098-1133. Patent English Entered STN: 9 Oct 2002 Last Updated on STN: 9 Oct 2002 The present invention relates to DNA sequences encoding ***Vmp*** -like polypeptides of pathogenic ***Borrelia*** , the use of the DNA sequences in recombinant vectors to express polypeptides, the encoded amino acid sequences, application of the DNA and amino acid sequences to the production of polypeptides as antigens for immunoprophylaxis, immunotherapy, and immunodiagnosis. Also disclosed are the use of the nucleic acid sequences as probes or primers for the deletion of organisms causing Lyme disease, relapsing fever, or related disorders, and kits designed to facilitate methods of using the described polypeptides, DNA segments and antibodies. ***VMP*** -like sequences of pathogenic ***borrelia*** ***Norris, Steven J.*** [Inventor, Reprint author]; Zhang, Jing-Ren [Inventor]; Hardham, John M. [Inventor]; Howell, Jerrilyn K. [Inventor]; Barbour, Alan G. [Inventor];. . The present invention relates to DNA sequences encoding ***Vmp*** -like ***Borrelia*** , the use of the DNA polypeptides of pathogenic sequences in recombinant vectors to express polypeptides, the encoded amino acid sequences, application of the. Major Concepts Infection; Molecular Genetics (Biochemistry and Molecular Biophysics) Chemicals & Biochemicals DNA sequences; ***Vmp*** -like polypeptides ORGN Classifier Spirochaetaceae 06112 Super Taxa Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name ***Borrelia*** : pathogen Taxa Notes Bacteria, Eubacteria, Microorganisms ANSWER 13 OF 21 CAPLUS COPYRIGHT 2009 ACS on STN 136:304895 ***Borrelia*** burgdorferi vlsE gene expression and Analysis of recombination in the tick vector Indest, Karl J.; Howell, Jerrilyn K.; Jacobs, Mary B.; Scholl-Meeker, Dorothy; ***Norris, Steven J.***; Philipp, Mario T. Department of Parasitology, Tulane Regional Primate Research Center, Tulane University Health Sciences Center, Covington, LA, 70433, USA Infection and Immunity (2001), 69(11), 7083-7090 CODEN: INFIBR; ISSN: 0019-9567

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Journal

American Society for Microbiology

- LA English
- AΒ Expression and recombination of the antigenic variation vlsE gene of the Lyme disease spirochete ***Borrelia*** burgdorferi were analyzed in the tick vector. To assess vlsE expression, Ixodes scapularis nymphs infected with the B. burgdorferi strain B31 were fed on mice for 48 or 96 h or to repletion, and then crushed and acetone fixed either immediately thereafter (ticks collected at the two earlier time points) or 4 days after repletion. Unfed nymphs also were examd. At all of the time points investigated, spirochetes were able to bind a rabbit antibody raised against the conserved invariable region 6 of VlsE, as assessed by indirect immunofluorescence, but not pre-immune serum from the same rabbit. This same antibody also bound to B31 spirochetes cultivated in vitro. Intensity of fluorescence appeared highest in cultured spirochetes, followed by spirochetes present in unfed ticks. Only a dim fluorescent signal was obsd. on spirochetes at the 48 and 96 h time points and at day 4 post-repletion. Expression of vlsE in vitro was affected by a rise in pH from 7.0 to 8.0 at 34.degree.C. Hence, vlsE expression appears to be sensitive to environmental cues of the type found in the B. burgdorferi natural history. To assess vlsE recombination, nymphs were capillary fed the B. burgdorferi B31 clonal isolate 5A3. Ticks thus infected were either left to rest for 4 wk (Group I) or fed to repletion on a mouse (Group II). The contents of each tick from both groups were cultured and 10 B. burgdorferi clones from the spirochetal isolate of each tick were obtained. The vlsE cassettes from several of these clones were amplified by PCR and sequenced. Regardless of whether the isolate was derived from Group I or Group II ticks, no changes were obsd. in the vlsE sequence. In contrast, vlsE cassettes amplified from B. burgdorferi clones derived from a mouse that was infected with B31-5A3 capillary-fed nymphs showed considerable recombination. It follows that vlsE recombination does not occur in the tick vector.
- RE.CNT 33 THERE ARE 33 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- TI Analysis of ***Borrelia*** burgdorferi vlsE gene expression and recombination in the tick vector
- AU Indest, Karl J.; Howell, Jerrilyn K.; Jacobs, Mary B.; Scholl-Meeker, Dorothy; ***Norris, Steven J.***; Philipp, Mario T.
- AB Expression and recombination of the antigenic variation vlsE gene of the Lyme disease spirochete ***Borrelia*** burgdorferi were analyzed in the tick vector. To assess vlsE expression, Ixodes scapularis nymphs infected with the B. burgdorferi strain. . .
- ST DNA sequence ***Borrelia*** gene vlsE mouse infection tick;

 Borrelia gene vlsE lipoprotein tick expression; protein sequence
 gene vlsE lipoprotein ***Borrelia***; genetic recombination

 Borrelia gene vlsE mouse infection tick
- IT Ixodes scapularis
 - (anal. of ***Borrelia*** burgdorferi vlsE gene expression and recombination in tick vector)
- IT Lipoproteins
 - RL: BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study)
 - (gene vlsE, for ***Vmp*** -like sequence; partial sequence and expression in tick of ***Borrelia*** burgdorferi gene vlsE lipoprotein)
- IT Development, nonmammalian postembryonic
 - (nymph; anal. of ***Borrelia*** burgdorferi vlsE gene expression
 and recombination in tick vector)
- IT ***Borrelia*** burgdorferi

DNA sequences

Protein sequences

(partial sequence of ***Borrelia*** burgdorferi gene vlsE lipoprotein isolated from mouse infected by infestation with Ixodes scapularis nymphal ticks)

IT Lyme disease

Mus

Recombination, genetic

(vlsE cassettes amplified from ***Borrelia*** burgdorferi clones derived from mouse infected with B31-5A3 capillary-fed nymphs showed considerable recombination)

IT Gene, microbial

RL: BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study)

(vlsE; partial DNA sequence, expression and recombination in tick
vector of ***Borrelia*** burgdorferi gene vlsE)

IT 411243-31-7 411243-32-8 411243-33-9 411243-34-0

RL: BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study)

(amino acid sequence; partial sequence of ***Borrelia***
burgdorferi gene vlsE lipoprotein isolated from mouse infected by
infestation with Ixodes scapularis nymphal ticks)

IT 359572-33-1, GenBank AY043397 359572-34-2, GenBank AY043398 359572-35-3, GenBank AY043399 359572-36-4, GenBank AY043400 382261-30-5, GenBank AY043401

RL: BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study)

(nucleotide sequence; partial DNA sequence, expression and recombination in tick vector of ***Borrelia*** burgdorferi gene vlsE)

- L2 ANSWER 14 OF 21 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN DUPLICATE 6
- AN 2001:63090 BIOSIS <<LOGINID::20090609>>
- DN PREV200100063090
- TI Correlation between plasmid content and infectivity in ***Borrelia*** burgdorferi.
- AU Purser, Joye E.; ***Norris, Steven J.*** [Reprint author]
- CS Department of Pathology and Laboratory Medicine, Medical School, and Graduate School of Biomedical, University of Texas-Houston Health Science Center, Houston, TX, 77225, USA Steven.J.Norris@uth.tmc.edu
- SO Proceedings of the National Academy of Sciences of the United States of America, (December 5, 2000) Vol. 97, No. 25, pp. 13865-13870. print. CODEN: PNASA6. ISSN: 0027-8424.
- DT Article
- LA English
- ED Entered STN: 31 Jan 2001 Last Updated on STN: 12 Feb 2002
- AB Infectivity-associated plasmids were identified in ***Borrelia*** burgdorferi B31 by using PCR to detect each of the plasmids in a panel of 19 clonal isolates. The clones exhibited high-, low-, and intermediate-infectivity phenotypes based on their frequency of isolation from needle-inoculated C3H/HeN mice. Presence or absence of 21 of the 22 plasmids was determined in each of the clones by using PCR primers specific for regions unique to each plasmid, as identified in the recently available genome sequence. Southern blot hybridization results were used

to confirm the PCR results in some cases. Plasmid 1p25 exhibited a direct correlation with infectivity in that it was consistently present in all clones of high or intermediate infectivity and was absent in all low-infectivity clones. lp28-1, containing the ***vmp*** -like sequence locus, also correlated with infectivity; all clones that lacked lp28-1 but contained 1p25 had an intermediate infectivity phenotype, in which infection was primarily restricted to the joints. Plasmids cp9, cp32-3, 1p21, 1p28-2, 1p28-4, and 1p56 apparently are not required for infection in this model, because clones lacking these plasmids exhibited a high-infectivity phenotype. Plasmids cp26, cp32-1, cp32-2 and/or cp32-7, cp32-4, cp32-6, cp32-8, cp32-9, lp17, lp28-3, lp36, lp38, and lp54 were consistently present in all clones examined. On the basis of these results, 1p25 and 1p28-1 appear to encode virulence factors important in the pathogenesis of B. burgdorferi B31.

- ΤI Correlation between plasmid content and infectivity in ***Borrelia*** burgdorferi.
- ΑU Purser, Joye E.; ***Norris, Steven J.*** [Reprint author]
- Infectivity-associated plasmids were identified in ***Borrelia*** AB burgdorferi B31 by using PCR to detect each of the plasmids in a panel of 19 clonal isolates. The clones. . . consistently present in all clones of high or intermediate infectivity and was absent in all low-infectivity clones. lp28-1, containing the ***vmp*** -like sequence locus, also correlated with infectivity; all clones that lacked 1p28-1 but contained 1p25 had an intermediate infectivity phenotype, in.

ORGN . . .

Mammals, Nonhuman Vertebrates, Nonhuman Mammals, Rodents, Vertebrates ORGN Classifier

Spirochaetaceae 06112

Super Taxa

Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name

Borrelia burgdorferi: pathogen

Bacteria, Eubacteria, Microorganisms

- ANSWER 15 OF 21 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on L2 DUPLICATE 7
- 2000:169153 BIOSIS <<LOGINID::20090609>> ΑN
- PREV200000169153 DN
- Conservation and heterogeneity of vlsE among human and tick isolates of TΙ ***Borrelia*** burgdorferi.
- Iyer, Radha; Hardham, John M.; Wormser, Gary P.; Schwartz, Ira; ΑU ***Norris, Steven J.*** [Reprint author]
- Department of Pathology and Laboratory Medicine, University of Texas CS Medical School at Houston, Houston, TX, 77225, USA
- SO Infection and Immunity, (March, 2000) Vol. 68, No. 3, pp. 1714-1718. CODEN: INFIBR. ISSN: 0019-9567.
- DT Article
- LA English
- EDEntered STN: 3 May 2000 Last Updated on STN: 4 Jan 2002
- The ***vls*** (variable major protein (***VMP***)-like sequence) AB locus of ***Borrelia*** burgdorferi encodes an antigenic variation system that closely resembles the ***VMP*** system of relapsing fever ***borreliae*** . To determine whether ***vls*** sequences are present consistently in low-passage, infectious isolates of B.

burgdorferi, 22 blood and erythema migrans biopsy isolates from Lyme disease patients in Westchester County, New York, were examined by Southern blot and PCR analysis. Each of the strains contained a single plasmid varying in size from 21 to 38 kb that hybridized strongly with a vlsE probe based on the B. burgdorferi B31 sequence. In contrast, PCR products were obtained with only 10 of the 22 strains when primers corresponding to the 5' and 3' regions of the B31 vlsE sequence outside the variable cassette region were used. Only 2 of 16 B. burgdorferi-infected tick specimens yielded detectable PCR product. Eight of 10 strains that yielded a PCR product under these conditions were type 1 (a genotype with a high rate of dissemination), according to PCR-restriction fragment length polymorphism analysis of intergenic rDNA sequences, whereas the isolates that did not yield vlsE PCR products were either type 2 or type 3. Comparison of the sequences of cloned PCR products from the patient isolates indicated a high degree of identity to the B31 sequence, with most of the differences restricted to the hypervariable regions known to undergo sequence variation. Taken together, these results both reinforce previous evidence that ***vls*** sequences are present consistently in low-passage Lyme disease spirochetes and indicate that both highly conserved and heterogeneous subgroups exist with regard to vlsE sequences.

TI Conservation and heterogeneity of vlsE among human and tick isolates of ***Borrelia*** burgdorferi.

AU Iyer, Radha; Hardham, John M.; Wormser, Gary P.; Schwartz, Ira; ***Norris, Steven J.*** [Reprint author]

AB The ***vls*** (variable major protein (***VMP***)-like sequence) locus of ***Borrelia*** burgdorferi encodes an antigenic variation system that closely resembles the ***VMP*** system of relapsing fever ***borreliae***. To determine whether ***vls*** sequences are present consistently in low-passage, infectious isolates of B. burgdorferi, 22 blood and erythema migrans biopsy isolates from Lyme. . . differences restricted to the hypervariable regions known to undergo sequence variation. Taken together, these results both reinforce previous evidence that ***vls*** sequences are present consistently in low-passage Lyme disease spirochetes and indicate that both highly conserved and heterogeneous subgroups exist with. . .

IT Major Concepts

Infection

IT Diseases

Lyme disease: bacterial disease

Lyme Disease (MeSH)

IT Chemicals & Biochemicals

Borrelia burgdorferi ***vls*** gene; ***Borrelia***

burgdorferi vlsE gene

ORGN . . .

Notes

Animals, Chordates, Humans, Mammals, Primates, Vertebrates $\operatorname{\mathsf{ORGN}}$ Classifier

Spirochaetaceae 06112

Super Taxa

Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name

Borrelia burgdorferi: pathogen

Taxa Notes

Bacteria, Eubacteria, Microorganisms

L2 ANSWER 16 OF 21 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on

STN DUPLICATE 8

- AN 1998:393355 BIOSIS <<LOGINID::20090609>>
- DN PREV199800393355
- TI Genetic variation of the ***Borrelia*** burgdorferi gene vlsE involves cassette-specific, segmental gene conversion.
- AU Zhang, Jing-Ren; ***Norris, Steven J.*** [Reprint author]
- CS Dep. Pathol. Lab. Med., Univ. Tex. Med. Sch., 6431 Fannin, Houston, TX 77030, USA
- SO Infection and Immunity, (Aug., 1998) Vol. 66, No. 8, pp. 3698-3704. print. CODEN: INFIBR. ISSN: 0019-9567.
- DT Article
- LA English
- ED Entered STN: 10 Sep 1998
 Last Updated on STN: 10 Sep 1998
- AΒ The Lyme disease spirochete ***Borrelia*** burgdorferi possesses 15 silent ***vls*** cassettes and a ***vls*** expression site (vlsE) encoding a surface-exposed lipoprotein. Segments of the silent ***vls*** cassettes have been shown to recombine with the vlsE cassette region in the mammalian host, resulting in combinatorial antigenic variation. Despite promiscuous recombination within the vlsE cassette region, the 5' and 3' coding sequences of vlsE that flank the cassette region are not subject to sequence variation during these recombination events. The segments of the silent ***vls*** cassettes recombine in the vlsE cassette region through a unidirectional process such that the sequence and organization of the silent ***vls*** loci are not affected. As a result of recombination, the previously expressed segments are replaced by incoming segments and apparently degraded. These results provide evidence for a gene conversion mechanism in VlsE antigenic variation.
- TI Genetic variation of the ***Borrelia*** burgdorferi gene vlsE involves cassette-specific, segmental gene conversion.
- AU Zhang, Jing-Ren; ***Norris, Steven J.*** [Reprint author]
- AB The Lyme disease spirochete ***Borrelia*** burgdorferi possesses 15 silent ***vls*** cassettes and a ***vls*** expression site (vlsE) encoding a surface-exposed lipoprotein. Segments of the silent ***vls*** cassettes have been shown to recombine with the vlsE cassette region in the mammalian host, resulting in combinatorial antigenic variation... that flank the cassette region are not subject to sequence variation during these recombination events. The segments of the silent ***vls*** cassettes recombine in the vlsE cassette region through a unidirectional process such that the sequence and organization of the silent ***vls*** loci are not affected. As a result of recombination, the previously expressed segments are replaced by incoming segments and apparently. . .

ORGN Classifier

Spirochaetaceae 06112

Super Taxa

Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name

Borrelia -burgdorferi

Taxa Notes

Bacteria, Eubacteria, Microorganisms

- L2 ANSWER 17 OF 21 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN DUPLICATE 9
- AN 1998:393354 BIOSIS <<LOGINID::20090609>>
- DN PREV199800393354

- TI Kinetics and in vivo induction of genetic variation of vlsE in ***Borrelia*** burgdorferi.
- AU Zhang, Jing-Ren; ***Norris, Steven J.*** [Reprint author]
- CS Dep. Pathol. Lab. Med., Univ. Tex. Med. Sch., 6431 Fannin, Houston, TX 77030, USA
- SO Infection and Immunity, (Aug., 1998) Vol. 66, No. 8, pp. 3689-3697. print. CODEN: INFIBR. ISSN: 0019-9567.
- DT Article
- LA English
- ED Entered STN: 10 Sep 1998

 Last Updated on STN: 10 Sep 1998
- AΒ The Lyme disease agent, ***Borrelia*** burgdorferi, is able to persistently infect humans and animals for months or years in the presence of an active immune response. It is not known how the organisms survive immune attack in the mammalian host. vlsE, a gene localized near one end of linear plasmid lp28-1 and encoding a surface-exposed lipoprotein in B. burgdorferi B31, was shown recently to undergo extensive genetic and antigenic variation within 28 days of initial infection in C3H/HeN mice. In this study, we examined the kinetics of vlsE sequence variation in C3H/HeN mice at 4, 7, 14, 21, and 28 days and at 7 and 12 months postinfection. Sequence changes were detected by PCR amplification and sequence analysis as early as 4 days postinfection and accumulated progressively in both C3H/HeN and CB-17 severe combined immunodeficient (SCID) mice throughout the course of infection. The sequence changes were consistent with sequential recombination of segments from multiple silent ***vls*** cassette sites into the vlsE expression site. No vlsE sequence changes were detected in organisms cultured in vitro for up to 84 days. These results indicate that vlsE recombination is induced by a factor(s) present in the mammalian host, independent of adaptive immune responses. The possible inducing conditions appear to be present in various tissue sites because isolates from multiple tissues showed similar degrees of sequence variation. The rate of accumulation of predicted amino acid changes was higher in the immunologically intact C3H/HeN mice than in SCID mice, a finding consistent with immune selection of VlsE variants.
- TI Kinetics and in vivo induction of genetic variation of vlsE in ***Borrelia*** burgdorferi.
- AU Zhang, Jing-Ren; ***Norris, Steven J.*** [Reprint author]
- AB The Lyme disease agent, ***Borrelia*** burgdorferi, is able to persistently infect humans and animals for months or years in the presence of an active immune. . . (SCID) mice throughout the course of infection. The sequence changes were consistent with sequential recombination of segments from multiple silent ***vls*** cassette sites into the vlsE expression site. No vlsE sequence changes were detected in organisms cultured in vitro for up. . .

ORGN . . .

 ${\tt Mammals,\ Nonhuman\ Vertebrates,\ Nonhuman\ Mammals,\ Rodents,\ Vertebrates}$ ${\tt ORGN\ Classifier}$

Spirochaetaceae 06112

Super Taxa

Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name

Borrelia -burgdorferi

Taxa Notes

Bacteria, Eubacteria, Microorganisms

L2 ANSWER 18 OF 21 CAPLUS COPYRIGHT 2009 ACS on STN

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1997:579836 CAPLUS <<LOGINID::20090609>>
ΑN
DN
    127:189742
OREF 127:36809a,36812a
      ***Vmp*** -like sequences of pathogenic ***Borrelia***
ΤI
IN
      ***Norris, Steven J.*** ; Zhang, Jing-ren; Hardham, John M.; Howell,
    Jerrilyn K.; Barbour, Alan G.; Weinstock, George M.
PΑ
    Board of Regents, the University of Texas System, USA; Norris, Steven J.;
    Zhang, Jing-Ren; Hardham, John M.; Howell, Jerrilyn K.; Barbour, Alan G.;
    Weinstock, George M.
SO
    PCT Int. Appl., 130 pp.
    CODEN: PIXXD2
DT
    Patent
    English
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    PATENT NO.
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    WO 9731123
                       A1 19970828 WO 1997-US2952 19970220
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            LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT,
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    US 2004-852555
                       А3
                            20040524
    The present invention relates to DNA sequences encoding ***Vmp*** -like
AΒ
    polypeptides of pathogenic ***Borrelia*** , the use of the DNA
    sequences in recombinant vectors to express polypeptides, the encoded
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amino acid sequences, application of the DNA and amino acid sequences to

immunotherapy, and immunodiagnosis. Also disclosed are the use of the

the prodn. of polypeptides as antigens for immunoprophylaxis,

nucleic acid sequences as probes or primers for the detection of organisms causing Lyme disease, relapsing fever, or related disorders, and kits designed to facilitate methods of using the described polypeptides, DNA segments and antibodies.

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THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT 1
             ALL CITATIONS AVAILABLE IN THE RE FORMAT
ΤI
       ***Vmp*** -like sequences of pathogenic ***Borrelia***
ΤN
       ***Norris, Steven J.*** ; Zhang, Jing-ren; Hardham, John M.; Howell,
     Jerrilyn K.; Barbour, Alan G.; Weinstock, George M.
AΒ
     The present invention relates to DNA sequences encoding
                                                            ***Vmp*** -like
    polypeptides of pathogenic ***Borrelia*** , the use of the DNA
     sequences in recombinant vectors to express polypeptides, the encoded
     amino acid sequences, application of the. .
ST
    variable major protein gene
                                  ***Borrelia***
ΙT
       ***Borrelia*** burgdorferi
        ( ***Vmp*** -like sequences of pathogenic ***Borrelia*** )
ΙT
    Proteins, specific or class
     RL: BOC (Biological occurrence); BSU (Biological study, unclassified);
     BIOL (Biological study); OCCU (Occurrence)
        (gene vlsE; ***Vmp*** -like sequences of pathogenic ***Borrelia***
       )
    189614-97-9, DNA ( ***Borrelia*** burgdorferi strain B31 clone 5A3 gene
ΤT
    vlsE plus flanks)
    RL: BOC (Biological occurrence); BSU (Biological study, unclassified);
    BIOL (Biological study); OCCU (Occurrence)
       ( ***Vmp*** -like sequences of pathogenic ***Borrelia*** )
ΙT
    189833-73-6, Protein ( ***Borrelia*** burgdorferi strain B31 clone 5A3
     gene vlsE)
     RL: BOC (Biological occurrence); BSU (Biological study, unclassified); THU
     (Therapeutic use); BIOL (Biological study); OCCU (Occurrence); USES (Uses)
        ( ***Vmp*** -like sequences of pathogenic
                                                    ***Borrelia*** )
    ANSWER 19 OF 21 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on
L2
                                                       DUPLICATE 10
    1997:225972 BIOSIS <<LOGINID::20090609>>
ΑN
    PREV199799517688
DN
ΤI
    Antigenic variation in Lyme disease ***Borreliae***
                                                            by promiscuous
    recombination of ***VMP*** -like sequence cassettes.
    Zhang, Jing-Ren [Reprint author]; Hardham, John M.; Barbour, Alan G.;
ΑU
      ***Norris, Steven J.***
CS
     Dep. Pathol., Univ. Texas Med. Sch. Houston, Houston, TX 77030, USA
    Cell, (1997) Vol. 89, No. 2, pp. 275-285.
SO
    CODEN: CELLB5. ISSN: 0092-8674.
DT
    Article
LA
    English
ED
    Entered STN: 22 May 1997
     Last Updated on STN: 22 May 1997
    We have identified and characterized an elaborate genetic system in the
AΒ
    Lyme disease spirochete ***Borrelia*** burgdorferi that promotes
     extensive antigenic variation of a surface-exposed lipoprotein, VlsE. A
     28 kb linear plasmid of B. burgdorferi B31 (lp28-1) was found to contain a
      ***vmp*** -like sequence ( ***vls*** ) locus that closely resembles
the
    variable major protein ( ***vmp*** ) system for antigenic variation of
    relapsing fever organisms. Portions of several of the 15 nonexpressed
     (silent) ***vls*** cassette sequences located upstream of vlsE
```

recombined into the central vlsE cassette region during infection of

C3H/HeN mice, resulting in antigenic variation of the expressed lipoprotein. This combinatorial variation could potentially produce millions of antigenic variants in the mammalian host.

- TI Antigenic variation in Lyme disease ***Borreliae*** by promiscuous recombination of ***VMP*** -like sequence cassettes.
- AU Zhang, Jing-Ren [Reprint author]; Hardham, John M.; Barbour, Alan G.;
 Norris, Steven J.
- AB We have identified and characterized an elaborate genetic system in the Lyme disease spirochete ***Borrelia*** burgdorferi that promotes extensive antigenic variation of a surface-exposed lipoprotein, VlsE. A 28 kb linear plasmid of B. burgdorferi B31 (lp28-1) was found to contain a ***vmp*** -like sequence (***vls***) locus that closely resembles

the variable major protein (***vmp***) system for antigenic variation of relapsing fever organisms. Portions of several of the 15 nonexpressed (silent) ***vls*** cassette sequences located upstream of vlsE recombined into the central vlsE cassette region during infection of C3H/HeN mice, resulting in. . .

IT Miscellaneous Descriptors

ANTIGENIC VARIATION; BACTERIAL DISEASE; COMBINATORIAL VARIATION; C3H/HEN; INFECTION; LYME DISEASE; MOLECULAR GENETICS; PATHOGEN; PROMISCUOUS RECOMBINATION; SURFACE-EXPOSED LIPOPROTEIN; VLSE; ***VMP*** -LIKE SEQUENCE CASSETTES

ORGN . .

Mammals, Nonhuman Vertebrates, Nonhuman Mammals, Rodents, Vertebrates ORGN Classifier

Spirochaetaceae 06112

Super Taxa

Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name

Borrelia burgdorferi

Taxa Notes

Bacteria, Eubacteria, Microorganisms

- L2 ANSWER 20 OF 21 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN
- AN 1997:282165 BIOSIS <<LOGINID::20090609>>
- DN PREV199799581368
- TI Antigenic variation in Lyme disease spirochetes by promiscuous recombination of ***vmp*** -like sequence cassettes.
- AU Zhang, Jing-Ren [Reprint author]; Hardham, John M.; Barbour, Alan G.; ***Norris, Steven J.***
- CS Dep. Pathol. Lab. Med., Univ. Texas Med. Sch., Houston, TX, USA
- SO Abstracts of the General Meeting of the American Society for Microbiology, (1997) Vol. 97, No. 0, pp. 103.

Meeting Info.: 97th General Meeting of the American Society for Microbiology. Miami Beach, Florida, USA. May 4-8, 1997. ISSN: 1060-2011.

- DT Conference; (Meeting)
 Conference; Abstract; (Meeting Abstract)
 Conference; (Meeting Poster)
- LA English
- ED Entered STN: 3 Jul 1997 Last Updated on STN: 3 Jul 1997
- TI Antigenic variation in Lyme disease spirochetes by promiscuous recombination of ***vmp*** -like sequence cassettes.
- AU Zhang, Jing-Ren [Reprint author]; Hardham, John M.; Barbour, Alan G.;

Norris, Steven J. ORGN . . . Mammals, Nonhuman Vertebrates, Nonhuman Mammals, Rodents, Vertebrates ORGN Classifier Spirochaetaceae 06112 Super Taxa Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name ***Borrelia*** burgdorferi Taxa Notes Bacteria, Eubacteria, Microorganisms L2 ANSWER 21 OF 21 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on DUPLICATE 11 ΑN 1994:345519 BIOSIS <<LOGINID::20090609>> PREV199497358519 DΝ ΤI A family of surface-exposed proteins of 20 kilodaltons in the genus ***Borrelia*** . Carter, Carol J.; Bergstrom, Sven; ***Norris, Steven J.***; Barbour, ΑU Alan G. [Reprint author] Dep. Microbiol. and Med., Univ. Texas Health Sci. Cent., San Antonio, TX CS 78284-7758, USA Infection and Immunity, (1994) Vol. 62, No. 7, pp. 2792-2799. SO CODEN: INFIBR. ISSN: 0019-9567. DT Article English LA OS Genbank-L24911 ΕD Entered STN: 8 Aug 1994 Last Updated on STN: 1 Sep 1994 Relapsing fever and Lyme disease spirochetes of the genus ***Borrelia*** AΒ display at their surfaces abundant lipoproteins: ***Vmp*** proteins in ***Borrelia*** hermsii and Osp proteins in ***Borrelia*** burgdorferi. ***Vmp*** and Osp proteins largely determine serotype specificity, and neutralizing antibodies of infected or immunized animals are directed at them. For the present study, we examined B. hermsii serotype 33, which is unique among strain HS1 serotypes in the low frequency of switches to other serotypes during infections and in vitro

cultivation. Failing to clone the complete ***vmp33*** gene, we accomplished its further characterization by (i) determining three partial amino acid sequences, (ii) designing oligonucleotide primers based on these amino acid sequences, (iii) cloning and sequencing the central portion of ***vmp33*** , and (iv) using outwardly directed primers and the inverse PCR to clone the 5' and 3' ends of the gene and flanking regions. The transcriptional start site was identified by primer extension analysis. ***Vmp33*** was a polypeptide of 211 amino acids; the three partial amino acid sequences were identified in the open reading ***Vmp33*** was found to be more similar to other 20-kDa ***Vmp*** proteins of B. hermsii and to OspC proteins of B. burgdorferi than t was to 35- to 39-kDa ***Vmp*** proteins of the same strain. Moreover, OspC proteins were more similar to ***Vmp33*** than they were to OspA, -B, or -D proteins of B. burgdorferi. These sequence similarities were consistent with Western blot (immunoblot) findings of crossreactions between ***Vmp33*** and OspC with anti- ***Vmp33*** and anti-OspC sera. The promoter for the expressed ***vmp33*** was found to be different from the expression site for other active ***vmp*** genes characterized to date. These results indicate that ***Vmp33*** and other small ***Vmp*** 's belong with OspC to a

```
genus-wide family of 20-kDa proteins and that expression of these proteins
     may be coordinated with expression of other ***Vmp*** and Osp proteins
     in ***Borrelia*** spp.
ΤI
    A family of surface-exposed proteins of 20 kilodaltons in the genus
      ***Borrelia*** .
     Carter, Carol J.; Bergstrom, Sven; ***Norris, Steven J.***; Barbour,
ΑU
     Alan G. [Reprint author]
AB
     Relapsing fever and Lyme disease spirochetes of the genus ***Borrelia***
     display at their surfaces abundant lipoproteins: ***Vmp*** proteins in
      ***Borrelia*** hermsii and Osp proteins in ***Borrelia***
     burgdorferi. ***Vmp*** and Osp proteins largely determine serotype
     specificity, and neutralizing antibodies of infected or immunized animals
     are directed at them. For. . . in the low frequency of switches to
     other serotypes during infections and in vitro cultivation. Failing to
     clone the complete ***vmp33*** gene, we accomplished its further
     characterization by (i) determining three partial amino acid sequences,
     (ii) designing oligonucleotide primers based on these amino acid
     sequences, (iii) cloning and sequencing the central portion of
      ***vmp33*** , and (iv) using outwardly directed primers and the inverse
     PCR to clone the 5' and 3' ends of the gene and flanking regions. The
     transcriptional start site was identified by primer extension analysis.
      ***Vmp33*** was a polypeptide of 211 amino acids; the three partial
     amino acid sequences were identified in the open reading frame.
      ***Vmp33*** was found to be more similar to other 20-kDa
     proteins of B. hermsii and to OspC proteins of B. burgdorferi than t was
     to 35- to 39-kDa ***Vmp*** proteins of the same strain. Moreover,
     OspC proteins were more similar to ***Vmp33*** than they were to OspA,
     -B, or -D proteins of B. burgdorferi. These sequence similarities were
     consistent with Western blot (immunoblot) findings of crossreactions
     between ***Vmp33*** and OspC with anti- ***Vmp33*** and anti-OspC
     sera. The promoter for the expressed ***vmp33*** gene was found to be
     different from the expression site for other active ***vmp*** genes
     characterized to date. These results indicate that ***Vmp33***
     other small ***Vmp*** 's belong with OspC to a genus-wide family of
     20-kDa proteins and that expression of these proteins may be coordinated
     with expression of other ***Vmp*** and Osp proteins in
      ***Borrelia***
                       spp.
IT
       sequence data; nucleotide sequence; L24911: Genbank
ΙT
    Miscellaneous Descriptors
       CLONING STRATEGY; HOMOLOGY; METHOD; OSPC PROTEIN; PROMOTER ANALYSIS;
       TRANSCRIPTION START SITE; ***VMP33*** GENE
ORGN Classifier
       Spirochaetaceae 06112
     Super Taxa
       Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms
     Organism Name
            ***Borrelia*** burgdorferi
            ***Borrelia*** hermsii
     Taxa Notes
       Bacteria, Eubacteria, Microorganisms
=> s borreli? and (VMP? or vls)
          337 BORRELI? AND (VMP? OR VLS)
=> dup rem 13
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L4 87 DUP REM L3 (250 DUPLICATES REMOVED)

=> d bib ab kwic 1-

YOU HAVE REQUESTED DATA FROM 87 ANSWERS - CONTINUE? Y/(N):y

- L4 ANSWER 1 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN DUPLICATE 1
- AN 2009:232172 BIOSIS <<LOGINID::20090609>>
- DN PREV200900232172
- TI Detailed Analysis of Sequence Changes Occurring during vlsE Antigenic Variation in the Mouse Model of ***Borrelia*** burgdorferi Infection.
- AU Coutte, Loic [Reprint Author]; Botkin, Douglas J.; Gao, Lihui; Norris, Steven J.
- CS Inst Biol Lille, Lille, France Steven.J.Norris@uth.tmc.edu
- SO PLoS Pathogens, (FEB 2009) Vol. 5, No. 2, pp. Article No.: e1000293. http://www.plospathogens.org.
 ISSN: 1553-7366. E-ISSN: 1553-7374.
- DT Article
- LA English
- ED Entered STN: 1 Apr 2009 Last Updated on STN: 1 Apr 2009
- Lyme disease ***Borrelia*** can infect humans and animals for months AΒ to years, despite the presence of an active host immune response. ***vls*** antigenic variation system, which expresses the surface-exposed lipoprotein VlsE, plays a major role in B. burgdorferi immune evasion. Gene conversion between ***vls*** silent cassettes and the vlsE expression site occurs at high frequency during mammalian infection, resulting in sequence variation in the VlsE product. In this study, we examined vlsE sequence variation in B. burgdorferi B31 during mouse infection by analyzing 1,399 clones isolated from bladder, heart, joint, ear, and skin tissues of mice infected for 4 to 365 days. The median number of codon changes increased progressively in C3H/HeN mice from 4 to 28 days post infection, and no clones retained the parental vlsE sequence at 28 days. In contrast, the decrease in the number of clones with the parental vlsE sequence and the increase in the number of sequence changes occurred more gradually in severe combined immunodeficiency (SCID) mice. Clones containing a stop codon were isolated, indicating that continuous expression of full-length VlsE is not required for survival in vivo; also, these clones continued to undergo vlsE recombination. Analysis of clones with apparent single recombination events indicated that recombinations into vlsE are nonselective with regard to the silent cassette utilized, as well as the length and location of the recombination event. Sequence changes as small as one base pair were common. Fifteen percent of recovered vlsE variants contained "template-independent'' sequence changes, which clustered in the variable regions of vlsE. We hypothesize that the increased frequency and complexity of vlsE sequence changes observed in clones recovered from immunocompetent mice (as compared with SCID mice) is due to rapid clearance of relatively invariant clones by variable region-specific anti-VlsE antibody responses.
- TI Detailed Analysis of Sequence Changes Occurring during vlsE Antigenic Variation in the Mouse Model of ***Borrelia*** burgdorferi Infection.
- AB Lyme disease ***Borrelia*** can infect humans and animals for months to years, despite the presence of an active host immune response. The ***vls*** antigenic variation system, which expresses the surface-exposed lipoprotein VlsE, plays a major role in B. burgdorferi

immune evasion. Gene conversion between ***vls*** silent cassettes and the vlsE expression site occurs at high frequency during mammalian infection, resulting in sequence variation in the. . .

IT . . .

system; bladder: excretory system; ear: sensory system; joint: skeletal system; skin tissue: integumentary system

IT Diseases

Lyme disease: bacterial disease, ***Borrelia*** burgdorferi infection

IT Chemicals & Biochemicals

antibody

ORGN . .

Mammals, Nonhuman Vertebrates, Nonhuman Mammals, Rodents, Vertebrates ORGN Classifier

Spirochaetaceae 06112

Super Taxa

Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name

Borrelia burgdorferi (species): pathogen, strain-B31 Taxa Notes

Bacteria, Eubacteria, Microorganisms

- L4 ANSWER 2 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN DUPLICATE 2
- AN 2008:2678 BIOSIS <<LOGINID::20090609>>
- DN PREV200800010948
- TI The role of VlsE antigenic variation in the Lyme disease spirochete: persistence through a mechanism that differs from other pathogens.
- AU Bankhead, Troy; Chaconas, George [Reprint Author]
- CS Univ Calgary, Dept Biochem and Mol Biol, Calgary, AB T2N 4N1, Canada chaconas@ucalgary.ca
- SO Molecular Microbiology, (SEP 2007) Vol. 65, No. 6, pp. 1547-1558. CODEN: MOMIEE. ISSN: 0950-382X.
- DT Article
- LA English

not.

- ED Entered STN: 12 Dec 2007 Last Updated on STN: 12 Dec 2007
- AB The linear plasmid, lp28-1, is required for persistent infection by the Lyme disease spirochete, ***Borrelia*** burgdorferi. This plasmid contains the ***vls*** antigenic variation locus, which has long been thought to be important for immune evasion. However, the role of the ***vls*** locus as a virulence factor during mammalian infection has

been clearly defined. We report the successful removal of the ***vls*** locus through telomere resolvase-mediated targeted deletion, and demonstrate the absolute requirement of this 1p28-1 component for persistence in the mouse host. Moreover, successful infection of C3H/HeN mice with an 1p28-1 plasmid in which the left portion was deleted excludes participation of other 1p28-1 non- ***vls*** genes in spirochete virulence, persistence and the process of recombinational switching at vlsE. Data are also presented that cast doubt on an immune evasion mechanism whereby VlsE directly masks other surface antigens similar to what has been observed for several other pathogens that undergo recombinational antigenic variation.

AB The linear plasmid, lp28-1, is required for persistent infection by the Lyme disease spirochete, ***Borrelia*** burgdorferi. This plasmid contains the ***vls*** antigenic variation locus, which has long been

thought to be important for immune evasion. However, the role of the ***vls*** locus as a virulence factor during mammalian infection has not. been clearly defined. We report the successful removal of the locus through telomere resolvase-mediated targeted deletion, and demonstrate the absolute requirement of this 1p28-1 component for persistence in the mouse. . . infection of C3H/HeN mice with an lp28-1plasmid in which the left portion was deleted excludes participation of other lp28-1 non- ***vls*** genes in spirochete virulence, persistence and the process of recombinational switching at vlsE. Data are also presented that cast doubt. ORGN . . . Mammals, Nonhuman Vertebrates, Nonhuman Mammals, Rodents, Vertebrates ORGN Classifier Spirochaetaceae 06112 Super Taxa Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name ***Borrelia*** burgdorferi (species): pathogen Bacteria, Eubacteria, Microorganisms ***Borrelia*** burgdorferi ***vls*** GEN gene (Spirochaetaceae) ANSWER 3 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN L42007:587055 BIOSIS <<LOGINID::20090609>> ΑN PREV200700592427 DN ΤI Gene conversion is a convergent strategy for pathogen antigenic variation. Palmer, Guy H. [Reprint Author]; Brayton, Kelly A. ΑU CS Washington State Univ, Programs Vector Borne Dis and Genom, Pullman, WA 99164 USA gpalmer@vetmed.wsu.edu Trends in Parasitology, (SEP 2007) Vol. 23, No. 9, pp. 408-413. SO ISSN: 1471-4922. DT Article General Review; (Literature Review) LA English Entered STN: 21 Nov 2007 EDLast Updated on STN: 21 Nov 2007 AΒ Recent studies on three unrelated vector-borne pathogens, Anaplasma marginale, ***Borrelia*** hermsii and Trypanosoma brucei, illustrate the central importance of gene conversion as a mechanism for antigenic variation, which results in subsequent evasion of the immune response and persistence in the reservoir host. The combination of genome sequence data and in vivo studies tracking variant emergence not only provides insight into the genetic mechanisms for variant generation and hierarchy in variant expression but also highlights gaps in our knowledge regarding variant capacity and usage in vivo. AB Recent studies on three unrelated vector-borne pathogens, Anaplasma ***Borrelia*** hermsii and Trypanosoma brucei, illustrate marginale, the central importance of gene conversion as a mechanism for antigenic variation, which results in. ΙT Population Genetics (Population Studies) ΙT Anaplasma marginale infection: parasitic disease ΙT Diseases

Trypanosoma brucei infection: parasitic disease

```
ΙT
    Diseases
           ***Borrelia*** hermsii infection: parasitic disease
ORGN . . .
       pathogen
     Taxa Notes
       Animals, Invertebrates, Microorganisms, Protozoans
ORGN Classifier
        Spirochaetaceae 06112
     Super Taxa
        Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms
     Organism Name
           ***Borrelia*** hermsii (species): pathogen
     Taxa Notes
       Bacteria, Eubacteria, Microorganisms
GEN
       ***Borrelia*** hermsii
                                 ***vmp***
                                             gene (Spirochaetaceae):
     expression; Trypanosoma brucei vsg gene (Flagellata): expression;
     Anaplasma marginale msp2 gene (Anaplasmataceae): expression
    ANSWER 4 OF 87 CABA COPYRIGHT 2009 CABI on STN DUPLICATE 3
L4
ΑN
     2007:261701 CABA <<LOGINID::20090609>>
     20073270507
DN
      ***Borrelia*** burgdorferi adhesins identified using in vivo phage
TΤ
    display
    Antonara, S.; Chafel, R. M.; LaFrance, M.; Coburn, J.
ΑU
    Graduate Program in Molecular Microbiology, Tufts University Sackler
CS
     School of Graduate Biomedical Sciences, Boston, Massachusetts, USA.
     jcoburn@tufts-nemc.org
SO
    Molecular Microbiology, (2007) Vol. 66, No. 1, pp. 262-276. many ref.
     Publisher: Blackwell Publishing. Oxford
     ISSN: 0950-382X
     URL: http://www.blackwell-synergy.com/loi/mmi
     DOI: 10.1111/j.1365-2958.2007.05924.x
CY
    United Kingdom
DT
    Journal
LA
    English
    Entered STN: 7 Dec 2007
ED
     Last Updated on STN: 7 Dec 2007
       ***Borrelia*** burgdorferi, the agent of Lyme disease, disseminates
AΒ
     from the site of deposition by Ixodes ticks to cause systemic infection.
     Dissemination occurs through the circulation and through tissue matrices,
     but the B. burgdorferi molecules that mediate interactions with the
     endothelium in vivo have not yet been identified. In vivo selection of
     filamentous phage expressing B. burgdorferi protein fragments on the phage
     surface identified several new candidate adhesins, and verified the
     activity of one adhesin that had been previously characterized in vitro.
     P66, a B. burgdorferi ligand for [beta]3-chain integrins, OspC, a protein
     that is essential for the establishment of infection in mammals, and
       ***Vls*** , a protein that undergoes antigenic variation in the mammal,
     were all selected for binding to the murine endothelium in vivo.
     Additional B. burgdorferi proteins for which no functions have been
     identified, including all four members of the OspF family and BmpD, were
     identified as candidate adhesins. The use of in vivo phage display is one
     approach to the identification of adhesins in pathogenic bacteria that are
     not easily grown in the laboratory, or for which genetic manipulations are
     not straightforward.
ΤI
       ***Borrelia*** burgdorferi adhesins identified using in vivo phage
```

display.

- ***Borrelia*** burgdorferi, the agent of Lyme disease, disseminates from the site of deposition by Ixodes ticks to cause systemic infection. Dissemination. . . B. burgdorferi ligand for [beta]3-chain integrins, OspC, a protein that is essential for the establishment of infection in mammals, and ***Vls*** , a protein that undergoes antigenic variation in the mammal, were all selected for binding to the murine endothelium in vivo.. .
- BT ***Borrelia*** ; Spirochaetaceae; Spirochaetales; Gracilicutes; bacteria; prokaryotes; Muridae; rodents; mammals; vertebrates; Chordata; animals; small mammals; eukaryotes
- ST lyme ***borreliosis***
- ORGN ***Borrelia*** burgdorferi; Murinae
- L4 ANSWER 5 OF 87 MEDLINE on STN
- AN 2006382551 MEDLINE <<LOGINID::20090609>>
- DN PubMed ID: 16796669
- TI Antigenic variation with a twist--the ***Borrelia*** story.
- AU Norris Steven J
- CS Department of Pathology. University of Texas Medical School at Houston, PO Box 20708, Houston, TX 77225-0708, USA.. Steven.J.Norris@uth.tmc.edu
- NC R01 AI37277 (United States NIAID NIH HHS)
- SO Molecular microbiology, (2006 Jun) Vol. 60, No. 6, pp. 1319-22. Journal code: 8712028. ISSN: 0950-382X.
- CY England: United Kingdom
- DT Commentary
 - Journal; Article; (JOURNAL ARTICLE)
 (RESEARCH SUPPORT, N.I.H., EXTRAMURAL)
- LA English
- FS Priority Journals
- EM 200608
- ED Entered STN: 27 Jun 2006

 Last Updated on STN: 23 Aug 2006

 Entered Medline: 22 Aug 2006
- A common mechanism of immune evasion in pathogenic bacteria and protozoa is antigenic variation, in which genetic or epigenetic changes result in rapid, sequential shifts in a surface-exposed antigen. In this issue of Molecular Microbiology, Dai et al. provide the most complete description to date of the vlp/vsp antigenic variation system of the relapsing fever spirochaete, ***Borrelia*** hermsii. This elaborate, plasmid-encoded system involves an expression site that can acquire either variable large protein (vlp) or variable small protein (vsp) surface lipoprotein genes from 59 different archival copies. The archival vlp and vsp genes are arranged in clusters on at least five different plasmids. Gene conversion occurs through recombination events at upstream homology sequences (UHS) found in each gene copy, and at downstream homology sequences (DHS) found periodically among the vlp/vsp archival genes. Previous studies have ***Borrelia*** shown that antigenic variation in relapsing fever only permits the evasion of host antibody responses, but can also result in changes in neurotropism and other pathogenic properties. The vlsE antigenic variation locus of Lyme disease spirochaetes, although similar in sequence to the relapsing fever vlp genes, has evolved a completely different antigenic variation mechanism involving segmental recombination from a contiguous array of ***vls*** silent cassettes. These two systems thus appear to represent divergence from a common precursor followed by functional convergence to create two distinct antigenic variation processes.
- TI Antigenic variation with a twist--the ***Borrelia*** story.

AB . . et al. provide the most complete description to date of the vlp/vsp antigenic variation system of the relapsing fever spirochaete, ***Borrelia*** hermsii. This elaborate, plasmid-encoded system involves an expression site that can acquire either variable large protein (vlp) or variable small. . . homology sequences (DHS) found periodically among the vlp/vsp archival genes. Previous studies have shown that antigenic variation in relapsing fever ***Borrelia*** not only permits the evasion of host antibody responses, but can also result in changes in neurotropism and other pathogenic. . . relapsing fever vlp genes, has evolved a completely different antigenic variation mechanism involving segmental recombination from a contiguous array of ***vls*** cassettes. These two systems thus appear to represent divergence from a common precursor followed by functional convergence to create. . . CT *Antigenic Variation: GE, genetics *Antigens, Bacterial: GE, genetics *** Borrelia: GE, genetics*** ****Borrelia: IM, immunology*** L4ANSWER 6 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN 2007:87547 BIOSIS <<LOGINID::20090609>> ΑN DN PREV200700093298 ***VMP*** -like sequences of pathogenic ***Borrelia*** ΤI Anonymous; Norris, Steven J. [Inventor]; Zhang, Jing-Ren [Inventor]; ΑU Hardham, John M. [Inventor]; Howell, Jerrilyn K. [Inventor]; Barbour, Alan G. [Inventor]; Weinstock, George M. [Inventor] CS Houston, TX USA ASSIGNEE: Board of Regents The University of Texas System PΙ US 07135176 20061114 Official Gazette of the United States Patent and Trademark Office Patents, SO (NOV 14 2006) CODEN: OGUPE7. ISSN: 0098-1133. DT Patent LA English ED Entered STN: 31 Jan 2007 Last Updated on STN: 31 Jan 2007 AΒ The present invention relates to DNA sequences encoding ***Vmp*** -like polypeptides of pathogenic ***Borrelia*** , the use of the DNA sequences in recombinant vectors to express polypeptides, the encoded amino acid sequences, application of the DNA and amino acid sequences to the production of polypeptides as antigens for immunoprophylaxis, immunotherapy, and immunodiagnosis. Also disclosed are the use of the nucleic acid sequences as probes or primers for the detection of organisms causing Lyme disease, relapsing fever, or related disorders, and kits designed to facilitate methods of using the described polypeptides, DNA segments and antibodies. ***VMP*** -like sequences of pathogenic ***Borrelia*** ΤI The present invention relates to DNA sequences encoding ***Vmp*** -like AΒ polypeptides of pathogenic ***Borrelia*** , the use of the DNA sequences in recombinant vectors to express polypeptides, the encoded amino acid sequences, application of the. ΙT Major Concepts Pharmacology; Clinical Immunology (Human Medicine, Medical Sciences); Infection TΤ Chemicals & Biochemicals ***VMP*** -like DNA sequences: diagnostic-drug, ***Borrelia*** immunostimulant-drug, immunologic-drug

- L4 ANSWER 7 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN DUPLICATE 4
- AN 2006:570967 BIOSIS <<LOGINID::20090609>>
- DN PREV200600576492
- TI Transcriptional regulation of the ***Borrelia*** burgdorferi antigenically variable VlsE surface protein.
- AU Bykowski, Tomasz; Babb, Kelly; von Lackum, Kate; Riley, Sean P.; Norris, Steven J.; Stevenson, Brian [Reprint Author]
- CS Univ Kentucky, Coll Med, Dept Microbiol Mol Genet and Immunol, Albert B Chandler Med Ctr, MS 415, Lexington, KY 40536 USA brian.stevenson@uky.edu
- SO Journal of Bacteriology, (JUL 2006) Vol. 188, No. 13, pp. 4879-4889. CODEN: JOBAAY. ISSN: 0021-9193.
- DT Article
- LA English
- ED Entered STN: 1 Nov 2006 Last Updated on STN: 1 Nov 2006
- The Lyme disease agent ***Borrelia*** burgdorferi can persistently AB infect humans and other animals despite host active immune responses. This is facilitated, in part, by the vis locus, a complex system consisting of the vlsE expression site and an adjacent set of 11 to 15 silent ***vls*** cassettes. Segments of nonexpressed cassettes recombine with the vlsE region during infection of mammalian hosts, resulting in combinatorial antigenic variation of the VISE outer surface protein. We now demonstrate that synthesis of VlsE is regulated during the natural mammal-tick infectious cycle, being activated in mammals but repressed during tick colonization. Examination of cultured B. burgdorferi cells indicated that the spirochete controls vlsE transcription levels in response to environmental cues. Analysis of PvlsE::gfp fusions in B. burgdorferi indicated that VlsE production is controlled at the level of transcriptional initiation, and regions of 5' DNA involved in the regulation were identified. Electrophoretic mobility shift assays detected qualitative and quantitative changes in patterns of protein-DNA complexes formed between the vlsE promoter and cytoplasmic proteins, suggesting the involvement of DNA-binding proteins in the regulation of vlsE, with at least one protein acting as a transcriptional activator.
- TI Transcriptional regulation of the ***Borrelia*** burgdorferi antigenically variable VlsE surface protein.
- AB The Lyme disease agent ***Borrelia*** burgdorferi can persistently infect humans and other animals despite host active immune responses. This is facilitated, in part, by the. . . vis locus, a complex system consisting of the vlsE expression site and an adjacent set of 11 to 15 silent ***vls*** cassettes. Segments of nonexpressed cassettes recombine with the vlsE region during infection of mammalian hosts, resulting in combinatorial antigenic variation. . .

Animals, Chordates, Humans, Mammals, Primates, Vertebrates ORGN Classifier

Spirochaetaceae 06112

Super Taxa

Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name

Borrelia burgdorferi (species): pathogen

Taxa Notes

Bacteria, Eubacteria, Microorganisms
Borrelia burgdorferi vlsE gene (Spirochaetaceae): expression

- L4 ANSWER 8 OF 87 CABA COPYRIGHT 2009 CABI on STN DUPLICATE 5
- AN 2006:126465 CABA <<LOGINID::20090609>>

DOI: 10.1111/j.1365-2958.2006.05204.x

DN 20063098478

GEN

- TI Antigenic variation with a twist the ***Borrelia*** story
- AU Norris, S. J.
- CS Department of Pathology & Laboratory Medicine, University of Texas Medical School at Houston, PO Box 20708, Houston, TX 77225-0708, USA. Steven.J.Norris@uth.tmc.edu
- SO Molecular Microbiology, (2006) Vol. 60, No. 6, pp. 1319-1322. 20 ref. Publisher: Blackwell Publishing. Oxford ISSN: 0950-382X URL: http://www.blackwell-synergy.com/servlet/useragent?func=showIssues&code=mmi
- CY United Kingdom
- DT Journal
- LA English
- ED Entered STN: 3 Aug 2006 Last Updated on STN: 3 Aug 2006
- A common mechanism of immune evasion in pathogenic bacteria and protozoa AΒ is antigenic variation, in which genetic or epigenetic changes result in rapid, sequential shifts in a surface-exposed antigen. In this issue of Molecular Microbiology, Dai et al. provide the most complete description to date of the vlp/vsp antigenic variation system of the relapsing fever ***Borrelia*** hermsii. This elaborate, plasmid-encoded spirochaete, system involves an expression site that can acquire either variable large protein (vlp) or variable small protein (vsp) surface lipoprotein genes from 59 different archival copies. The archival vlp and vsp genes are arranged in clusters on at least five different plasmids. Gene conversion occurs through recombination events at upstream homology sequences (UHS) found in each gene copy, and at downstream homology sequences (DHS) found periodically among the vlp/vsp archival genes. Previous studies have shown that antigenic variation in relapsing fever ***Borrelia*** not only permits the evasion of host antibody responses, but can also result in changes in neurotropism and other pathogenic properties. The vlsE antigenic variation locus of Lyme disease spirochaetes, although similar in sequence to the relapsing fever vlp genes, has evolved a completely different antigenic variation mechanism involving segmental recombination ***vls*** silent cassettes. These two from a contiquous array of systems thus appear to represent divergence from a common precursor followed by functional convergence to create two distinct antigenic variation processes.
- TI Antigenic variation with a twist the ***Borrelia*** story.
- AB . . . et al. provide the most complete description to date of the vlp/vsp antigenic variation system of the relapsing fever spirochaete, ***Borrelia*** hermsii. This elaborate, plasmid-encoded system involves an expression site that can acquire either variable large protein (vlp) or variable small. . . homology sequences (DHS) found periodically among the vlp/vsp archival genes. Previous studies have shown that antigenic variation in relapsing fever ***Borrelia*** not only permits the evasion of host antibody responses, but can also result in changes in neurotropism and other pathogenic. . . relapsing fever vlp genes, has evolved a completely different antigenic variation mechanism involving segmental recombination from a contiguous array of ***vls*** silent

cassettes. These two systems thus appear to represent divergence from a common precursor followed by functional convergence to create. . . ВΤ Spirochaetaceae; Spirochaetales; Gracilicutes; bacteria; prokaryotes; ***Borrelia*** ***Borrelia*** ; ***Borrelia*** ORGN hermsii L4ANSWER 9 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN DUPLICATE 6 ΑN 2006:406251 BIOSIS <<LOGINID::20090609>> DN PREV200600407284 ΤI Immunodominant epitope in the C-terminus of a variable major protein in ***Borrelia*** duttonii, an agent of tick-borne relapsing fever. ΑU Tabuchi, Norihiko [Reprint Author]; Tomoda, Koichiro; Kawaguchi, Hiroshi; Iwamoto, Hiroyuki; Fukunaga, Masahito CS Fukuyama Univ, Fac Pharm and Pharmaceut Sci, Mol Microbiol Lab, Gakuencho 1, Hiroshima 7290292, Japan tabuchi@fupharm.fukuyama-u.ac.jp SO Microbiology and Immunology, (2006) Vol. 50, No. 4, pp. 293-305. CODEN: MIIMDV. ISSN: 0385-5600. DT Article LA English Entered STN: 17 Aug 2006 ED Last Updated on STN: 17 Aug 2006 ***Borrelia*** duttonii strain Ly was isolated from a child with AΒ tick-borne relapsing fever in Tanzania. B. duttonii produces variable major proteins (***Vmps***), which undergo antigenic variation. We previously reported transcription of the ***vmpP*** gene, which is one ***Vmp*** genes in strain Ly, detected in vitro cultivation. of the In the current study, we purified the recombinant non-lipidated ***VmpP*** protein by affinity chromatography and produced polyclonal antibodies. Antigenicity of ***VmpP*** was examined by Western immunoblot analysis and peptide-based enzyme-linked immunosorbent assays. Antigenic epitopes were shown to comprise five regions interspersed within the ***VmpP*** primary amino acid sequence. Synthetic peptides spanning residues of three of five regions, 232-237 (LASIVD), 280-285 (AGGIAL), and 350-355 (KAADQQ), reacted strongly with the ***VmpP*** -specific antibody and these residues were identified as epitopes. In particular, the C-terminal domain (KAADQQ) of this protein was immunoreactive. Further research based on our results will promote the development of a recombinant vaccine for B. duttonii infection. TΙ Immunodominant epitope in the C-terminus of a variable major protein in ***Borrelia*** duttonii, an agent of tick-borne relapsing fever. ***Borrelia*** duttonii strain Ly was isolated from a child with AB tick-borne relapsing fever in Tanzania. B. duttonii produces variable major proteins (***Vmps***), which undergo antigenic variation. We previously reported transcription of the ***vmpP*** gene, which is one of the ***Vmp*** genes in strain Ly, detected in vitro cultivation. In the current study, we purified the recombinant non-lipidated ***VmpP*** protein by affinity chromatography and produced polyclonal antibodies. Antigenicity of ***VmpP*** was examined by Western immunoblot analysis and peptide-based enzyme-linked immunosorbent assays. Antigenic epitopes were shown to comprise five regions interspersed within the ***VmpP*** primary amino acid sequence. Synthetic peptides spanning residues of three of five regions, 232-237 (LASIVD), 280-285 (AGGIAL), and 350-355 (KAADQQ), reacted strongly with ***VmpP*** -specific antibody and these residues were identified as

epitopes. In particular, the C-terminal domain (KAADQQ) of this protein

```
was immunoreactive. Further. . .
ORGN . . .
Notes
       Animals, Chordates, Humans, Mammals, Primates, Vertebrates
ORGN Classifier
       Spirochaetaceae 06112
     Super Taxa
       Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms
     Organism Name
           ***Borrelia*** duttonii (species): pathogen
     Taxa Notes
       Bacteria, Eubacteria, Microorganisms
GEN
       ***Borrelia*** duttonii ***vmpP*** gene gene (Spirochaetaceae)
L4
    ANSWER 10 OF 87
                       MEDLINE on STN
                                                       DUPLICATE 7
    2006527807
                 MEDLINE <<LOGINID::20090609>>
ΑN
DN
    PubMed ID: 16914037
ΤI
    Comparative genome analysis: selection pressure on the
                                                             ***Borrelia***
                  cassettes is essential for infectivity.
ΑU
    Glockner Gernot; Schulte-Spechtel Ulrike; Schilhabel Markus; Felder
    Marius; Suhnel Jurgen; Wilske Bettina; Platzer Matthias
    Genome Analysis Group, Leibniz Institute for Age Research - Fritz Lipmann
CS
    Institute, Beutenbergstr, 11, 07745 Jena, Germany.. gernot@fli-leibniz.de
    BMC genomics, (2006) Vol. 7, pp. 211. Electronic Publication: 2006-08-16.
SO
    Journal code: 100965258. E-ISSN: 1471-2164.
    Report No.: NLM-PMC1559707.
CY
    England: United Kingdom
DT
    (COMPARATIVE STUDY)
    Journal; Article; (JOURNAL ARTICLE)
     (RESEARCH SUPPORT, NON-U.S. GOV'T)
    English
LA
    Priority Journals
FS
    200610
EM
ED
    Entered STN: 6 Sep 2006
    Last Updated on STN: 19 Oct 2006
    Entered Medline: 18 Oct 2006
AΒ
    BACKGROUND: At least three species of ***Borrelia*** burgdorferi sensu
    lato (Bbsl) cause tick-borne Lyme disease. Previous work including the
     genome analysis of B. burgdorferi B31 and B. garinii PBi suggested a
     highly variable plasmid part. The frequent occurrence of duplicated
     sequence stretches, the observed plasmid redundancy, as well as the mainly
    unknown function and variability of plasmid encoded genes rendered the
     relationships between plasmids within and between species largely
     unresolvable. RESULTS: To gain further insight into ***Borreliae***
     genome properties we completed the plasmid sequences of B. garinii PBi,
     added the genome of a further species, B. afzelii PKo, to our analysis,
     and compared for both species the genomes of pathogenic and apathogenic
     strains. The core of all Bbsl genomes consists of the chromosome and two
    plasmids collinear between all species. We also found additional groups
     of plasmids, which share large parts of their sequences. This makes it
     very likely that these plasmids are relatively stable and share common
     ancestors before the diversification of ***Borrelia***
                                                              species.The
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analysis of the differences between B. garinii PBi and B. afzelii PKo genomes of low and high passages revealed that the loss of infectivity is

Whereas B. garinii PBi suffered only from the break-off of a plasmid end, B. afzelii PKo lost more material, probably an entire plasmid. In both

accompanied in both species by a loss of similar genetic material.

cases the ***vls*** gene locus encoding for variable surface proteins is affected. CONCLUSION: The complete genome sequences of a B. garinii and a B. afzelii strain facilitate further comparative studies within the genus Borrellia. Our study shows that loss of infectivity can be traced back to only one single event in B. garinii PBi: the loss of the ***vls*** cassettes possibly due to error prone gene conversion. Similar albeit extended losses in B. afzelii PKo support the hypothesis that infectivity of ***Borrelia*** species depends heavily on the evasion from the host response. Comparative genome analysis: selection pressure on the ***Borrelia*** ***vls*** cassettes is essential for infectivity. BACKGROUND: At least three species of ***Borrelia*** burgdorferi sensu lato (Bbsl) cause tick-borne Lyme disease. Previous work including the genome analysis of B. burgdorferi B31 and B.. . plasmid encoded genes rendered the relationships between plasmids within and between species largely unresolvable. RESULTS: To gain further insight into ***Borreliae*** genome properties we completed the plasmid sequences of B. garinii PBi, added the genome of a further species, B. afzelii. . . sequences. This makes it very likely that these plasmids are relatively stable and share common ancestors before the diversification of ***Borrelia*** species. The analysis of the differences between B. garinii PBi and B. afzelii PKo genomes of low and high passages revealed. . . the break-off of a plasmid end, B. afzelii PKo lost more material, probably an entire plasmid. In both cases the ***vls*** gene locus encoding for variable surface proteins is affected. CONCLUSION: The complete genome sequences of a B. garinii and a. . . loss of infectivity can be traced back to only one single event in B. garinii PBi: ***vls*** the loss of the cassettes possibly due to error prone gene conversion. Similar albeit extended losses in B. afzelii PKo support the hypothesis that infectivity of ***Borrelia*** species depends heavily on the evasion from the host response. ****Borrelia: GE, genetics*** *** Borrelia: PY, pathogenicity*** *** Borrelia Infections: MI, microbiology*** *** Borrelia burgdorferi: GE, genetics*** *** Borrelia burgdorferi: PY, pathogenicity*** Chromosomes, Bacterial: GE, genetics DNA, Bacterial: CH, chemistry DNA, Bacterial: GE, genetics Genes, Bacterial: GE, genetics ANSWER 11 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on DUPLICATE 8 2006:675157 BIOSIS <<LOGINID::20090609>>

- L4
- ΑN
- DM PREV200600667335

ΤI

AB

CT

- Comparative genome analysis: selection pressure on the ***Borrelia*** ΤI ***vls*** cassettes is essential for infectivity.
- ΑU Gloeckner, Gernot [Reprint Author]; Schulte-Spechtel, Ulrike; Schilhabel, Markus; Felder, Marius; Suehnel, Juergen; Wilske, Bettina; Platzer, Matthias
- CS Fritz Lipmann Inst, Leibniz Inst Age Res, Genome Anal Grp, Beutenbergstr 11, D-07745 Jena, Germany gernot@fli-leibniz.de; Spechtel@m3401.mpk.med.uni-muenchen.de; mbs@fli-leibniz.de; mfelder@fli-leibniz.de; jsuehnel@fli-leibniz.de; Bettina.Wilske@mvp-bak.med.uni-muenchen.de; mplatzer@fli-leibniz.de
- BMC Genomics, (AUG 16 2006) Vol. 7, pp. Article No.: 211. SO

ISSN: 1471-2164.

- DT Article
- LA English
- OS genBank-CP000397CP000406
- ED Entered STN: 6 Dec 2006
 - Last Updated on STN: 20 Sep 2007
- AB Background: At least three species of ***Borrelia*** burgdorferi sensu lato (Bbsl) cause tick-borne Lyme disease. Previous work including the genome analysis of B. burgdorferi B31 and B. garinii PBi suggested a highly variable plasmid part. The frequent occurrence of duplicated sequence stretches, the observed plasmid redundancy, as well as the mainly unknown function and variability of plasmid encoded genes rendered the relationships between plasmids within and between species largely unresolvable.Results: To gain further insight into ***Borreliae*** genome properties we completed the plasmid sequences of B. garinii PBi, added the genome of a further species, B. afzelii PKo, to our analysis, and compared for both species the genomes of pathogenic and apathogenic strains. The core of all Bbsl genomes consists of the chromosome and two plasmids collinear between all species. We also found additional groups of plasmids, which share large parts of their sequences. This makes it very likely that these plasmids are relatively stable and share common ancestors before the diversification of ***Borrelia*** species.The analysis of the differences between B. garinii PBi and B. afzelii PKo genomes of low and high passages revealed that the loss of infectivity is accompanied in both species by a loss of similar genetic material. Whereas B. garinii PBi suffered only from the break-off of a plasmid end, B. afzelii PKo lost more material, probably an entire plasmid. In both ***vls*** cases the gene locus encoding for variable surface proteins is affected. Conclusion: The complete genome sequences of a B. garinii and a B. afzelii strain facilitate further comparative studies within the genus Borrellia. Our study shows that loss of infectivity can be traced back to only one single event in B. garinii PBi: the loss of the ***vls*** cassettes possibly due to error prone gene conversion. Similar albeit extended losses in B. afzelii PKo support the hypothesis ***Borrelia*** that infectivity of species depends heavily on the evasion from the host response.
- TI Comparative genome analysis: selection pressure on the ***Borrelia***

 vls cassettes is essential for infectivity.
- AB Background: At least three species of ***Borrelia*** burgdorferi sensu lato (Bbsl) cause tick-borne Lyme disease. Previous work including the genome analysis of B. burgdorferi B31 and B. . . of plasmid encoded genes rendered the relationships between plasmids within and between species largely unresolvable.Results: To gain further insight into
 - ***Borreliae*** genome properties we completed the plasmid sequences of B. garinii PBi, added the genome of a further species, B. afzelii. . . sequences. This makes it very likely that these plasmids are relatively stable and share common ancestors before the diversification of
 - ***Borrelia*** species.The analysis of the differences between B. garinii PBi and B. afzelii PKo genomes of low and high passages revealed.

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on the evasion from the host response. ORGN . . . Notes Animals, Chordates, Humans, Mammals, Primates, Vertebrates ORGN Classifier Spirochaetaceae 06112 Super Taxa Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name ***Borrelia*** burgdorferi (species): pathogen, strain-B31 garinii (species): pathogen, strain-PBi ***Borrelia*** ***Borrelia*** afzelii (species): pathogen, strain-PKo Taxa Notes Bacteria, Eubacteria, Microorganisms L4ANSWER 12 OF 87 CABA COPYRIGHT 2009 CABI on STN DUPLICATE 9 AN 2007:2067 CABA <<LOGINID::20090609>> DN 20063185553 ΤI Comparative genome analysis: selection pressure on the ***Borrelia*** ***vls*** cassettes is essential for infectivity ΑU Glockner, G.; Schulte-Spechtel, U.; Schilhabel, M.; Felder, M.; Suhnel, J.; Wilske, B.; Platzer, M. Genome Analysis Group, Leibniz Institute for Age Research-Fritz Lipmann CS Institute, Beutenbergstr. 11, 07745 Jena, Germany. gernot@fli-leibniz.de; Spechtel@m3401.mpk.med.uni-muenchen.de; mbs@fli-leibniz.de; mfelder@fli-leibniz.de; jsuehnel@fli-leibniz.de; Bettina.Wilske@mvp-bak.med.uni-muenchen.de; mplatzer@fli-leibniz.de SO BMC Genomics, (2006) Vol. 7, No. 211, pp. (16 August 2006). 36 ref. Publisher: BioMed Central Ltd. London ISSN: 1471-2164 URL: http://www.biomedcentral.com/content/pdf/1471-2164-7-211.pdf CY United Kingdom Journal DT LA English ED Entered STN: 8 Jan 2007 Last Updated on STN: 8 Jan 2007 AΒ Background: At least 3 species of ***Borrelia*** burgdorferi sensu lato (Bbsl) cause tickborne Lyme disease. Previous work, including the genome analysis of B. burgdorferi B31 and B. garinii PBi, suggested a highly variable plasmid part. The frequent occurrence of duplicated sequence stretches, the observed plasmid redundancy, as well as the mainly unknown function and variability of plasmid encoded genes render the relationships between plasmids within and between species largely unresolvable. Results: To gain further insight into the ***Borrelia*** genome properties, the plasmid sequences of B. garinii PBi were completed, the genome of B. afzelii PKo was analysed, and the genomes of the pathogenic and apathogenic strains of both species were compared. The core of all Bbsl genomes consisted of the chromosome and 2 plasmids collinear between all species. Additional groups of plasmids, which share large parts of their sequences, were observed, suggesting that these plasmids are relatively stable and share common ancestors before the

diversification of ***Borrelia*** species. The analysis of the

lost more material, probably an entire plasmid. In both cases, the

differences between B. garinii PBi and B. afzelii PKo genomes of low and high passages revealed that the loss of infectivity was accompanied in both species by a loss of similar genetic material. B. garinii PBi

suffered only from the break-off of a plasmid end, whereas B. afzelii PKo

vls gene locus encoding for variable surface proteins was affected. Conclusion: The complete genome sequences of a B. garinii and a B. afzelii strain facilitate further comparative studies within the genus ***Borrelia*** . The loss of infectivity can be traced back to only one single event in B. garinii PBi: the loss of the ***vls*** cassettes is possibly due to error prone gene conversion. Similar albeit extended losses in B. afzelii PKo support the hypothesis that the infectivity of ***Borrelia*** species depends heavily on the evasion from the host response.

TI Comparative genome analysis: selection pressure on the ***Borrelia***

vls cassettes is essential for infectivity.

Background: At least 3 species of ***Borrelia*** burgdorferi sensu AB lato (Bbsl) cause tickborne Lyme disease. Previous work, including the genome analysis of B. burgdorferi B31 and B.. . . encoded genes render the relationships between plasmids within and between species largely unresolvable. Results: To gain further insight into the ***Borrelia*** genome properties, the plasmid sequences of B. garinii PBi were completed, the genome of B. afzelii PKo was analysed, and. . . of their sequences, were observed, suggesting that these plasmids are relatively stable and share common ancestors before the diversification of ***Borrelia*** species. The analysis of the differences between B. garinii PBi and B. afzelii PKo genomes of low and high passages. . . break-off of a plasmid end, whereas B. afzelii PKo lost more material, probably an entire plasmid. In both cases, the ***vls*** gene locus encoding for variable surface proteins was affected. Conclusion: The complete genome sequences of a B. garinii and a B. afzelii strain facilitate further comparative studies within the genus ***Borrelia*** . The loss of infectivity can be traced back to only one single event in B. garinii PBi: the loss of the ***vls*** cassettes is possibly due to error prone gene conversion. Similar albeit extended losses in B. afzelii PKo support the hypothesis ***Borrelia*** species depends heavily on the that the infectivity of evasion from the host response.

BT ***Borrelia*** ; Spirochaetaceae; Spirochaetales; Gracilicutes; bacteria; prokaryotes

ORGN ***Borrelia*** afzelii; ***Borrelia*** burgdorferi; ***Borrelia*** garinii

- L4 ANSWER 13 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN
- AN 2006:134820 BIOSIS <<LOGINID::20090609>>
- DN PREV200600145254
- TI ***Vmp*** -like sequences of pathogenic ***Borrelia***
- AU Norris, Steven J. [Inventor]; Zhang, Jing-Ren [Inventor]; Hardham, John M. [Inventor]; Howell, Jerrilyn K. [Inventor]; Barbour, Alan G. [Inventor]; Weinstock, George M. [Inventor]
- CS Houston, TX USA
 - ASSIGNEE: Board of Regents, The University of Texas System
- PI US 06878816 20050412
- SO Official Gazette of the United States Patent and Trademark Office Patents, (APR 12 2005)
 CODEN: OGUPE7. ISSN: 0098-1133.
- DT Patent
- LA English
- ED Entered STN: 22 Feb 2006 Last Updated on STN: 22 Feb 2006
- AB The present invention relates to DNA sequences encoding ***Vmp*** -like polypeptides of pathogenic ***Borrelia*** , the use of the DNA

sequences in recombinant vectors to express polypeptides, the encoded amino acid sequences, application of the DNA and amino acid sequences to the production of polypeptides as antigens for immunoprophylaxis, immunotherapy, and immunodiagnosis. Also disclosed are the use of the nucleic acid sequences as probes or primers for the detection of organisms causing Lyme disease, relapsing fever, or related disorders, and kits designed to facilitate methods of using the described polypeptides, DNA segments and antibodies.

TI ***Vmp*** -like sequences of pathogenic ***Borrelia*** .

AB The present invention relates to DNA sequences encoding ***Vmp*** -like polypeptides of pathogenic ***Borrelia*** , the use of the DNA sequences in recombinant vectors to express polypeptides, the encoded amino acid sequences, application of the. . .

IT . . .

Clinical Immunology (Human Medicine, Medical Sciences); Infection; Clinical Chemistry (Allied Medical Sciences); Molecular Genetics (Biochemistry and Molecular Biophysics)

IT Diseases

Borrelia infection: bacterial disease, drug therapy, prevention and control

IT Chemicals & Biochemicals

DNA sequences; ***Vmp*** -like sequences; ***Borrelia*** polypeptide antigens: diagnostic-drug, immunostimulant-drug, immunologic-drug

ORGN Classifier

Spirochaetaceae 06112

Super Taxa

Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name

Borrelia (genus): pathogen

Taxa Notes

Bacteria, Eubacteria, Microorganisms

- L4 ANSWER 14 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN DUPLICATE 10
- AN 2005:554633 BIOSIS <<LOGINID::20090609>>
- DN PREV200510340099
- TI Variable tick protein in two genomic groups of the relapsing fever spirochete ***Borrelia*** hermsii in western North America.
- AU Porcella, Stephen F.; Raffel, Sandra J.; Anderson, Donald E. Jr; Gilk, Stacey D.; Bono, James L.; Schrumpf, Merry E.; Schwan, Tom G. [Reprint Author]
- CS NIAID, Rocky Mt Labs, Lab Human Bacterial Pathogenesis, 903 S 4th St, Hamilton, MT 59840 USA tom_schwan@nih.gov
- SO Infection and Immunity, (OCT 2005) Vol. 73, No. 10, pp. 6647-6658. CODEN: INFIBR. ISSN: 0019-9567.
- DT Article
- LA English
- ED Entered STN: 7 Dec 2005 Last Updated on STN: 7 Dec 2005
- AB ***Borrelia*** hermsii is the primary cause of tick-borne relapsing fever in North America. When its tick vector, Ornithodoros hermsi, acquires these spirochetes from the blood of an infected mammal, the bacteria switch their outer surface from one of, many bloodstream variable major proteins (***Vmps***) to a unique protein, Vtp (Vsp33). Vtp may be critical for successful tick transmission of B. hermsii; however, the

gene encoding this protein has been described previously in only one isolate. Here we identified and sequenced the vtp gene in 31 isolates of B. hermsii collected over 40 years from localities throughout much of its known geographic distribution. Seven major Vtp types were found. Little or no sequence variation existed within types, but between them significant variation was observed,, similar to the pattern of diversity described for the outer surface protein C (OspC) gene in Lyme disease spirochetes. The pattern of sequence relatedness among the Vtp types was incongruent in two branches compared to two genomic groups identified among the isolates by multilocus sequence typing of the 16S rRNA, flaB, gyrB, and glpQ genes. Therefore, both horizontal transfer and recombination within and between the two genomic, groups were responsible for some of the variation observed in the vtp gene. O. hermsi ticks were capable of transmitting spirochetes in the newly identified genomic group. Therefore, given the longevity of the tick vector and persistent infection of spirochetes in ticks, these arthropods rather than mammals may be the likely host where the exchange of spirochetal DNA occurs.

TI Variable tick protein in two genomic groups of the relapsing fever spirochete ***Borrelia*** hermsii in western North America.

AB ***Borrelia*** hermsii is the primary cause of tick-borne relapsing fever in North America. When its tick vector, Ornithodoros hermsi, acquires these. . . from the blood of an infected mammal, the bacteria switch their outer surface from one of, many bloodstream variable major proteins (***Vmps***) to a unique protein, Vtp (Vsp33). Vtp may be critical for successful tick transmission of B. hermsii; however, the gene. . .

ORGN . .

Mammals, Nonhuman Vertebrates, Nonhuman Mammals, Rodents, Vertebrates ORGN Classifier

Spirochaetaceae 06112

Super Taxa

Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name

Borrelia hermsii (species): pathogen

Taxa Notes

Bacteria, Eubacteria, Microorganisms

- GEN ***Borrelia*** hermsii vtp gene (Spirochaetaceae); ***Borrelia*** hermsii flaB gene (Spirochaetaceae); ***Borrelia*** hermsii glpQ gene (Spirochaetaceae)
- L4 ANSWER 15 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN
- AN 2004:283015 BIOSIS <<LOGINID::20090609>>
- DN PREV200400283530
- TI ***VMP*** -like sequences of pathogenic ***borrelia***
- AU Norris, Steven J. [Inventor, Reprint Author]; Zhang, Jing-Ren [Inventor]; Hardham, John M. [Inventor]; Howell, Jerrilyn K. [Inventor]; Barbour, Alan G. [Inventor]; Weinstock, George M. [Inventor]
- CS ASSIGNEE: Board of Regents, The University of Texas System
- PI US 6740744 20040525
- SO Official Gazette of the United States Patent and Trademark Office Patents, (May 25 2004) Vol. 1282, No. 4.
 - http://www.uspto.gov/web/menu/patdata.html. e-file.
 ISSN: 0098-1133 (ISSN print).
- DT Patent
- LA English
- ED Entered STN: 9 Jun 2004

Last Updated on STN: 9 Jun 2004 AΒ The present invention relates to DNA sequences encoding ***Vmp*** -like polypeptides of pathogenic ***Borrelia*** , the use of the DNA sequences in recombinant vectors to express polypeptides, the encoded amino acid sequences, application of the DNA and amino acid sequences to the production of polypeptides as antigens for immunoprophylaxis, immunotherapy, and immunodiagnosis. Also disclosed are the use of the nucleic acid sequences as probes or primers for the detection of organisms causing Lyme disease, relapsing fever, or related disorders, and kits designed to facilitate methods of using the described polypeptides, DNA segments and antibodies. ***VMP*** -like sequences of pathogenic ***borrelia*** ΤI AΒ The present invention relates to DNA sequences encoding ***Vmp*** -like ***Borrelia*** , the use of the DNA polypeptides of pathogenic sequences in recombinant vectors to express polypeptides, the encoded amino acid sequences, application of the. . . ΙT Major Concepts Equipment Apparatus Devices and Instrumentation; Infection; Methods and Techniques; Molecular Genetics (Biochemistry and Molecular Biophysics) ΙT ***Borrelia*** infection: bacterial disease ***Borrelia*** Infections (MeSH) ΙT Diseases Lyme disease: bacterial disease, diagnosis Lyme Disease (MeSH) ΤT Diseases relapsing fever: bacterial disease, diagnosis Relapsing Fever (MeSH) Chemicals & Biochemicals ΤТ ***Vmp*** -like polypeptides: encoding DNA sequences, encoding amino acid sequences; antibodies ΤT Methods & Equipment ***Borrelia*** infection assay method: bioassay techniques, laboratory techniques; immunodiagnosis: immunologic techniques, laboratory techniques; immunoprophylaxis: immunologic techniques, laboratory techniques; immunotherapy: clinical techniques,. . . ORGN Classifier Spirochaetaceae 06112 Super Taxa Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name ***Borrelia*** (genus): pathogen Bacteria, Eubacteria, Microorganisms ANSWER 16 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on L4ΑN 2004:257493 BIOSIS <<LOGINID::20090609>> DN PREV200400257602 TΙ ***VMP*** -like sequences of pathogenic ***Borrelia*** Norris, Steven J. [Inventor, Reprint Author]; Zhang, Jing-Ren [Inventor]; ΑU Hardham, John M. [Inventor]; Howell, Jerrilyn K. [Inventor]; Barbour, Alan G. [Inventor]; Weinstock, George M. [Inventor] CS Delmar, NY, USA ASSIGNEE: Board of Regents, The University of Texas System PΙ US 6719983 20040413

SO Official Gazette of the United States Patent and Trademark Office Patents, (Apr 13 2004) Vol. 1281, No. 2. http://www.uspto.gov/web/menu/patdata.html. e-file. ISSN: 0098-1133 (ISSN print). DT Patent LA English ED Entered STN: 12 May 2004 Last Updated on STN: 12 May 2004 The present invention relates to DNA sequences encoding ***Vmp*** -like AΒ polypeptides of pathogenic ***Borrelia*** , the use of the DNA sequences in recombinant vectors to express polypeptides, the encoded amino acid sequences, application of the DNA and amino acid sequences to the production of polypeptides as antigens for immunoprophylaxis, immunotherapy, and immunodiagnosis. Also disclosed are the use of the nucleic acid sequences as probes or primers for the detection of organisms causing Lyme disease, relapsing fever, or related disorders, and kits designed to facilitate methods of using the described polypeptides, DNA segments and antibodies. ΤI ***VMP*** -like sequences of pathogenic ***Borrelia*** The present invention relates to DNA sequences encoding ***Vmp*** -like AΒ polypeptides of pathogenic ***Borrelia*** , the use of the DNA sequences in recombinant vectors to express polypeptides, the encoded amino acid sequences, application of the. ΙT Major Concepts Medical Genetics (Allied Medical Sciences); Molecular Genetics (Biochemistry and Molecular Biophysics) ΙT Chemicals & Biochemicals ***Borrelia*** ***VMP*** -like DNA sequences ANSWER 17 OF 87 CAPLUS COPYRIGHT 2009 ACS on STN L42004:1033546 CAPLUS <<LOGINID::20090609>> ΑN 142:22291 DN Nucleic acid and/or polypeptide sequences of ***Borrelia*** ΤI burgdorferi for vaccination and antibody preparation techniques Sykes, Kathryn F.; Hale, Katherine S.; Johnston, Stephen A. ΙN Macrogenics, Inc., USA; Board of Regents, the University of Texas System PΑ SO PCT Int. Appl., 121 pp. CODEN: PIXXD2 DTPatent LA English FAN.CNT 1 KIND DATE APPLICATION NO. DATE PATENT NO. -----____ _____ A2 20041202 A3 20051229 WO 2004103269 20041202 WO 2003-US33056 PΙ 20031017 WO 2004103269 W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR,

BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

A1 20041213 AU 2003-304145

A1 20050317 US 2003-688058

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PRAI US 2002-419401P
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     WO 2003-US33056 W
                                20031017
     The invention relates to 34 antigens and nucleic acids encoding such
     antigens obtainable by screening a ***Borrelia*** genome, in
     particular a B. burgdorferi genome. In more specific aspects, the
     invention relates to methods of isolating such antigens and nucleic acids
     and to methods of using such isolated antigens for producing immune
     responses. The ability of an antigen to produce an immune response may be
     employed in vaccination or antibody prepn. techniques.
             THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD
             ALL CITATIONS AVAILABLE IN THE RE FORMAT
     Nucleic acid and/or polypeptide sequences of
                                                   ***Borrelia***
TΙ
     burgdorferi for vaccination and antibody preparation techniques
AΒ
     The invention relates to 34 antigens and nucleic acids encoding such
     antigens obtainable by screening a ***Borrelia*** genome, in
     particular a B. burgdorferi genome. In more specific aspects, the
     invention relates to methods of isolating such antigens.
SYSTEM LIMIT EXCEEDED DURING KWIC/STRING SEARCH
     antigen gene sequence ***Borrelia***
                                             vaccine antibody
IT
     Gene, microbial
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (BB00043; nucleic acid and/or polypeptide sequences of ***Borrelia***
        burgdorferi for vaccination and antibody prepn. techniques)
     Gene, microbial
ΤT
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (BB0043; nucleic acid and/or polypeptide sequences of ***Borrelia***
        burgdorferi for vaccination and antibody prepn. techniques)
    Gene, microbial
ΙT
     Gene, microbial
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (BB0072; nucleic acid and/or polypeptide sequences of
                                                               ***Borrelia***
        burgdorferi for vaccination and antibody prepn. techniques)
ΙT
     Gene, microbial
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (BB0133; nucleic acid and/or polypeptide sequences of ***Borrelia***
        burgdorferi for vaccination and antibody prepn. techniques)
ΙT
     Gene, microbial
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (BB0351; nucleic acid and/or polypeptide sequences of ***Borrelia***
        burgdorferi for vaccination and antibody prepn. techniques)
ΤТ
     Gene, microbial
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (BB0451; nucleic acid and/or polypeptide sequences of ***Borrelia***
        burgdorferi for vaccination and antibody prepn. techniques)
ΙT
     Gene, microbial
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (BB0508; nucleic acid and/or polypeptide sequences of ***Borrelia***
        burgdorferi for vaccination and antibody prepn. techniques)
ΙT
     Gene, microbial
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
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use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (BB0540; nucleic acid and/or polypeptide sequences of ***Borrelia***
        burgdorferi for vaccination and antibody prepn. techniques)
ΙT
     Gene, microbial
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (BB0656; nucleic acid and/or polypeptide sequences of
        burgdorferi for vaccination and antibody prepn. techniques)
ΤТ
     Gene, microbial
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (BBA04; nucleic acid and/or polypeptide sequences of ***Borrelia***
        burgdorferi for vaccination and antibody prepn. techniques)
ΙT
     Gene, microbial
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (BBB14; nucleic acid and/or polypeptide sequences of
                                                               ***Borrelia***
        burgdorferi for vaccination and antibody prepn. techniques)
ΤТ
    Gene, microbial
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (BBE02; nucleic acid and/or polypeptide sequences of
                                                              ***Borrelia***
        burgdorferi for vaccination and antibody prepn. techniques)
ΙT
    Gene, microbial
    RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (BBF05; nucleic acid and/or polypeptide sequences of
        burgdorferi for vaccination and antibody prepn. techniques)
ΤТ
     Gene, microbial
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (BBF13; nucleic acid and/or polypeptide sequences of
                                                               ***Borrelia***
        burgdorferi for vaccination and antibody prepn. techniques)
ΙT
     Gene, microbial
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (BBG24; nucleic acid and/or polypeptide sequences of
        burgdorferi for vaccination and antibody prepn. techniques)
     Gene, microbial
ΤТ
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (BBJ12; nucleic acid and/or polypeptide sequences of
                                                               ***Borrelia***
        burgdorferi for vaccination and antibody prepn. techniques)
ΙT
     Gene, microbial
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (BBM10; nucleic acid and/or polypeptide sequences of
                                                               ***Borrelia***
        burgdorferi for vaccination and antibody prepn. techniques)
ΤТ
     Gene, microbial
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (BBM11; nucleic acid and/or polypeptide sequences of ***Borrelia***
        burgdorferi for vaccination and antibody prepn. techniques)
ΙT
     Gene, microbial
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (BBO11; nucleic acid and/or polypeptide sequences of ***Borrelia***
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burgdorferi for vaccination and antibody prepn. techniques)
ΙT
     Gene, microbial
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (BBO29; nucleic acid and/or polypeptide sequences of
                                                               ***Borrelia***
        burgdorferi for vaccination and antibody prepn. techniques)
ΙT
     Gene, microbial
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (BBR01; nucleic acid and/or polypeptide sequences of
                                                              ***Borrelia***
        burgdorferi for vaccination and antibody prepn. techniques)
     Gene, microbial
ΙT
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (BBS36; nucleic acid and/or polypeptide sequences of
                                                               ***Borrelia***
        burgdorferi for vaccination and antibody prepn. techniques)
ΙT
     Gene, microbial
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (BBT01; nucleic acid and/or polypeptide sequences of
        burgdorferi for vaccination and antibody prepn. techniques)
ΤT
     Plasmids
        (CP32-7; nucleic acid and/or polypeptide sequences of
                                                               ***Borrelia***
        burgdorferi for vaccination and antibody prepn. techniques)
     Translation elongation factors
ΤT
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (EF-G; nucleic acid and/or polypeptide sequences of
                                                              ***Borrelia***
        burgdorferi for vaccination and antibody prepn. techniques)
ΙT
     Proteins
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (GTP-binding; nucleic acid and/or polypeptide sequences of
          ***Borrelia*** burgdorferi for vaccination and antibody prepn.
        techniques)
     Gene, microbial
ΤT
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (PGK; nucleic acid and/or polypeptide sequences of
                                                            ***Borrelia***
        burgdorferi for vaccination and antibody prepn. techniques)
ΙT
    Antigens
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (S2; nucleic acid and/or polypeptide sequences of
                                                           ***Borrelia***
        burgdorferi for vaccination and antibody prepn. techniques)
ΤТ
     Proteins
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (Vls8; nucleic acid and/or polypeptide sequences of
                                                              ***Borrelia***
        burgdorferi for vaccination and antibody prepn. techniques)
ΙT
     Proteins
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (Vls9; nucleic acid and/or polypeptide sequences of ***Borrelia***
        burgdorferi for vaccination and antibody prepn. techniques)
ΙT
    Proteins
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
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use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (VlsE1; nucleic acid and/or polypeptide sequences of
                                                              ***Borrelia***
        burgdorferi for vaccination and antibody prepn. techniques)
ΙT
     Transport proteins
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (chromate-transporting; nucleic acid and/or polypeptide sequences of
          ***Borrelia***
                         burgdorferi for vaccination and antibody prepn.
        techniques)
ΙT
     Plasmids
        (cp26; nucleic acid and/or polypeptide sequences of
                                                              ***Borrelia***
        burgdorferi for vaccination and antibody prepn. techniques)
ΙT
        (cp32-3; nucleic acid and/or polypeptide sequences of
                                                                ***Borrelia***
        burgdorferi for vaccination and antibody prepn. techniques)
ΙT
        (cp32-4; nucleic acid and/or polypeptide sequences of
                                                                ***Borrelia***
        burgdorferi for vaccination and antibody prepn. techniques)
ΙT
     Plasmids
                                                                ***Borrelia***
        (cp32-6; nucleic acid and/or polypeptide sequences of
        burgdorferi for vaccination and antibody prepn. techniques)
ΙT
     Plasmids
        (cp32-7; nucleic acid and/or polypeptide sequences of
                                                                ***Borrelia***
        burgdorferi for vaccination and antibody prepn. techniques)
     Herpesviridae
ΤT
        (detection of; nucleic acid and/or polypeptide sequences of
          ***Borrelia*** burgdorferi for vaccination and antibody prepn.
        techniques)
ΙT
     Proteins
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (gene BB0043; nucleic acid and/or polypeptide sequences of
          ***Borrelia*** burgdorferi for vaccination and antibody prepn.
        techniques)
ΙT
    Proteins
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (gene BB007224; nucleic acid and/or polypeptide sequences of
          ***Borrelia*** burgdorferi for vaccination and antibody prepn.
        techniques)
ΙT
     Proteins
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (gene BB0072; nucleic acid and/or polypeptide sequences of
          ***Borrelia***
                          burgdorferi for vaccination and antibody prepn.
        techniques)
ΤТ
     Proteins
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (gene BB0133; nucleic acid and/or polypeptide sequences of
          ***Borrelia*** burgdorferi for vaccination and antibody prepn.
        techniques)
TΤ
     Proteins
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (gene BB0351; nucleic acid and/or polypeptide sequences of
          ***Borrelia*** burgdorferi for vaccination and antibody prepn.
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techniques)
ΙT
     Proteins
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (gene BB0451; nucleic acid and/or polypeptide sequences of
          ***Borrelia*** burgdorferi for vaccination and antibody prepn.
        techniques)
ΤТ
    Proteins
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (gene BBB14; nucleic acid and/or polypeptide sequences of
          ***Borrelia*** burgdorferi for vaccination and antibody prepn.
        techniques)
ΙT
    Proteins
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (gene BBE02; nucleic acid and/or polypeptide sequences of
          ***Borrelia*** burgdorferi for vaccination and antibody prepn.
        techniques)
ΙT
     Proteins
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (gene BBF05; nucleic acid and/or polypeptide sequences of
          ***Borrelia*** burgdorferi for vaccination and antibody prepn.
        techniques)
ΤT
     Proteins
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (gene BBF13; nucleic acid and/or polypeptide sequences of
          ***Borrelia*** burgdorferi for vaccination and antibody prepn.
        techniques)
ΤТ
    Proteins
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (gene BBG24; nucleic acid and/or polypeptide sequences of
          ***Borrelia*** burgdorferi for vaccination and antibody prepn.
        techniques)
ΙT
     Proteins
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (gene BBJ12; nucleic acid and/or polypeptide sequences of
          ***Borrelia*** burgdorferi for vaccination and antibody prepn.
        techniques)
ΙT
     Proteins
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (gene BBM10; nucleic acid and/or polypeptide sequences of
          ***Borrelia***
                         burgdorferi for vaccination and antibody prepn.
        techniques)
ΤТ
     Proteins
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (gene BBM11; nucleic acid and/or polypeptide sequences of
          ***Borrelia*** burgdorferi for vaccination and antibody prepn.
        techniques)
ΙT
    Proteins
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RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic

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use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (gene BB011; nucleic acid and/or polypeptide sequences of
          ***Borrelia*** burgdorferi for vaccination and antibody prepn.
        techniques)
ΤТ
     Proteins
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (gene BB029; nucleic acid and/or polypeptide sequences of
          ***Borrelia*** burgdorferi for vaccination and antibody prepn.
        techniques)
ΙT
     Proteins
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (gene BBR01; nucleic acid and/or polypeptide sequences of
          ***Borrelia*** burgdorferi for vaccination and antibody prepn.
        techniques)
ΙT
     Proteins
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (gene BBS36; nucleic acid and/or polypeptide sequences of
          ***Borrelia*** burgdorferi for vaccination and antibody prepn.
        techniques)
ΙT
    Proteins
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (gene BBT01; nucleic acid and/or polypeptide sequences of
          ***Borrelia*** burgdorferi for vaccination and antibody prepn.
        techniques)
ΙT
     Gene, microbial
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (glpK; nucleic acid and/or polypeptide sequences of
                                                              ***Borrelia***
        burgdorferi for vaccination and antibody prepn. techniques)
ΙT
     Gene, microbial
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (gluA; nucleic acid and/or polypeptide sequences of
                                                              ***Borrelia***
        burgdorferi for vaccination and antibody prepn. techniques)
ΤТ
     Transport proteins
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (glycine/betaine/proline-binding; nucleic acid and/or polypeptide
                      ***Borrelia*** burgdorferi for vaccination and
        sequences of
        antibody prepn. techniques)
ΙT
     Gene, microbial
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (gpsA; nucleic acid and/or polypeptide sequences of
        burgdorferi for vaccination and antibody prepn. techniques)
     Antibodies and Immunoglobulins
IT
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (humanized; nucleic acid and/or polypeptide sequences of
          ***Borrelia*** burgdorferi for vaccination and antibody prepn.
       techniques)
TΤ
        (1p25; nucleic acid and/or polypeptide sequences of
                                                              ***Borrelia***
        burgdorferi for vaccination and antibody prepn. techniques)
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ΙT
    Plasmids
     Plasmids
     Plasmids
        (1p28-1; nucleic acid and/or polypeptide sequences of
                                                                ***Borrelia***
        burgdorferi for vaccination and antibody prepn. techniques)
ΤТ
    Plasmids
        (1p28-2; nucleic acid and/or polypeptide sequences of
        burgdorferi for vaccination and antibody prepn. techniques)
ΤТ
     Plasmids
        (lp38; nucleic acid and/or polypeptide sequences of
                                                              ***Borrelia***
        burgdorferi for vaccination and antibody prepn. techniques)
ΙT
        (1p54; nucleic acid and/or polypeptide sequences of
                                                              ***Borrelia***
        burgdorferi for vaccination and antibody prepn. techniques)
ΤТ
     Plasmids
        (1p56; nucleic acid and/or polypeptide sequences of
                                                               ***Borrelia***
        burgdorferi for vaccination and antibody prepn. techniques)
ΙT
     Plasmids
                                                             ***Borrelia***
        (lp5; nucleic acid and/or polypeptide sequences of
        burgdorferi for vaccination and antibody prepn. techniques)
ΤТ
     Gene, microbial
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (mfd; nucleic acid and/or polypeptide sequences of
        burgdorferi for vaccination and antibody prepn. techniques)
ΤТ
    Gene, microbial
     Proteins
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (mutL; nucleic acid and/or polypeptide sequences of
                                                             ***Borrelia***
        burgdorferi for vaccination and antibody prepn. techniques)
ΤТ
    Aptamers
         ***Borrelia***
         ***Borrelia***
                        burgdorferi
     DNA sequences
     Immunoassay
     Lyme disease
     Molecular cloning
     Protein sequences
     Vaccines
        (nucleic acid and/or polypeptide sequences of ***Borrelia***
        burgdorferi for vaccination and antibody prepn. techniques)
    Antibodies and Immunoglobulins
ΤТ
     RL: ANT (Analyte); ARG (Analytical reagent use); BPN (Biosynthetic
     preparation); DGN (Diagnostic use); NUU (Other use, unclassified); ANST
     (Analytical study); BIOL (Biological study); PREP (Preparation); USES
     (Uses)
        (nucleic acid and/or polypeptide sequences of
        burgdorferi for vaccination and antibody prepn. techniques)
IT
    Antigens
     Gene, microbial
     Proteins
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); PRP
     (Properties); THU (Therapeutic use); BIOL (Biological study); PREP
     (Preparation); USES (Uses)
        (nucleic acid and/or polypeptide sequences of ***Borrelia***
        burgdorferi for vaccination and antibody prepn. techniques)
```

```
Gene, microbial
TТ
    RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (proX; nucleic acid and/or polypeptide sequences of
                                                             ***Borrelia***
       burgdorferi for vaccination and antibody prepn. techniques)
ΙT
    Gene, microbial
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (rho; nucleic acid and/or polypeptide sequences of
                                                           ***Borrelia***
        burgdorferi for vaccination and antibody prepn. techniques)
ΙT
    Proteins
    RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (transcription-repair coupling factor; nucleic acid and/or polypeptide
        sequences of
                     ***Borrelia*** burgdorferi for vaccination and
       antibody prepn. techniques)
ΙT
    Gene, microbial
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (trxB; nucleic acid and/or polypeptide sequences of
       burgdorferi for vaccination and antibody prepn. techniques)
ΤT
    Immunization
        (vaccination; nucleic acid and/or polypeptide sequences of
          ***Borrelia*** burgdorferi for vaccination and antibody prepn.
       techniques)
ΤT
    Gene, microbial
    Gene, microbial
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
                                                            ***Borrelia***
        (vls8; nucleic acid and/or polypeptide sequences of
       burgdorferi for vaccination and antibody prepn. techniques)
ΙT
    Gene, microbial
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (vlsE1; nucleic acid and/or polypeptide sequences of
                                                              ***Borrelia***
       burgdorferi for vaccination and antibody prepn. techniques)
ΙT
     800367-85-5P
                   800367-86-6P 800367-87-7P
                                                800926-61-8P, Protein
     (plasmid lp56 fragment)
                              800926-63-0P, Protein (plasmid cp32-4 gene
             800926-65-2P
                           800926-67-4P
                                          800926-69-6P
                                                         800926-71-0P
     BBR01)
     800926-73-2P
                  800926-75-4P, Protein (plasmid lp25 gene BBE02)
     800926-77-6P 800926-79-8P, Protein (plasmid cp32-7 gene BB011)
     800926-81-2P 800926-83-4P, Protein (plasmid lp28-1 gene BBF13)
     800926-85-6P
                   800926-87-8P
                                  800926-89-0P
                                                 800926-91-4P
                                                                800926-93-6P
                                  800926-99-2P, Protein (plasmid cp32-7 gene
     800926-95-8P
                   800926-97-0P
     BB029)
             800927-02-0P
                           800927-04-2P
                                          800927-06-4P
                                                         800927-08-6P
     800927-10-0P, Protein (plasmid 1p28-2 gene BBG24)
                                                      800927-12-2P
     800927-14-4P
                   800927-16-6P
                                  800927-18-8P
                                                 800927-20-2P
                                                                800927-22-4P,
     Protein (plasmid cp32-6 gene BBM11)
                                          800927-24-6P 800927-26-8P
     800927-28-0P
                  800927-30-4P, Protein (plasmid lp28-1 gene BBF05)
                   800927-34-8P, Protein (plasmid cp32-6 gene BBM10)
     800927-32-6P
     800927-36-0P
                   800927-38-2P, Protein (plasmid cp32-3 gene BBS36)
     800927-40-6P 800927-42-8P 800927-44-0P
                                                800927-46-2P
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                                  800927-54-2P
     800927-50-8P 800927-52-0P
                                                 800927-56-4P
                                                                800927-58-6P
     800927-60-0P, Protein (plasmid cp26 gene BBB14)
                                                     800927-62-2P
     800927-64-4P 800927-66-6P 800927-68-8P
                                                800927-70-2P
                                                                800927-73-5P,
    Protein (plasmid lp28-1 gene vlsE1) 800927-75-7P
                                                         800927-77-9P
     800927-79-1P 800927-82-6P 800927-84-8P 800927-86-0P, Protein
```

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(plasmid lp5 gene BBT01) 800927-89-3P 800927-91-7P 800927-93-9P
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (amino acid sequence; nucleic acid and/or polypeptide sequences of
         ***Borrelia*** burgdorferi for vaccination and antibody prepn.
       techniques)
    182909-40-6, GenBank U43414 200798-27-2, GenBank AE000793
    GenBank AE000785 200798-31-8, GenBank AE000784 200798-32-9, GenBank
    AE000789 200798-33-0, GenBank AE000788 200798-34-1, GenBank AE000787
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     247563-26-4, GenBank AE001576 247563-27-5, GenBank AE001577
     247563-28-6, GenBank AE001578 247563-29-7, GenBank AE001579
     247563-30-0, GenBank AE001580 247563-31-1, GenBank AE001581
     247563-32-2, GenBank AE001582 247563-33-3, GenBank AE001584
     254953-60-1, GenBank AF169008
     RL: BSU (Biological study, unclassified); PRP (Properties); BIOL
     (Biological study)
        (nucleic acid and/or polypeptide sequences of ***Borrelia***
       burgdorferi for vaccination and antibody prepn. technique)
IT
    9001-83-6P, Phosphoglycerate kinase 9030-66-4P, Glycerol kinase
     9074-14-0P, Thioredoxin reductase 9075-65-4P, Glycerol-3-phosphate
    dehydrogenase 9076-84-0P, Coproporphyrinogen III oxidase 295324-05-9P,
     Glutamyl-tRNA amidotransferase
    RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
       (nucleic acid and/or polypeptide sequences of ***Borrelia***
       burgdorferi for vaccination and antibody prepn. techniques)
ΙT
    800926-60-7P, DNA (plasmid lp56 gene fragment) 800926-62-9P, DNA
     (plasmid cp32-4 gene BBR01)
                                800926-64-1P
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     800926-74-3P, DNA (plasmid 1p25 gene BBE02) 800926-76-5P, DNA (plasmid
     cp32-7 gene BB011 fragment) 800926-78-7P, DNA (plasmid cp32-7 gene
             800926-80-1P
     BB011)
   800926-82-3P, DNA (plasmid lp28-1 gene BBF13) 800926-84-5P
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                                                               800926-96-9P,
     DNA (plasmid cp32-7 gene BB029 fragment) 800926-98-1P, DNA (plasmid
     cp32-7 gene BBO29) 800927-00-8P, DNA (plasmid lp38 gene BBJ12 fragment)
     800927-01-9P, DNA (plasmid lp38 gene BBJ12) 800927-03-1P
                                                              800927-05-3P
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     (plasmid lp28-2 gene BBG24)
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     800927-17-7P
                   800927-19-9P, DNA (plasmid cp32-6 gene BBM11 fragment)
     800927-21-3P, DNA (plasmid cp32-6 gene BBM11)
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     800927-25-7P
                   800927-27-9P, DNA (plasmid lp28-1 gene BBF05 fragment)
     800927-29-1P, DNA (plasmid 1p28-1 gene BBF05) 800927-31-5P, DNA (plasmid
     cp32-6 gene BBM10 fragment) 800927-33-7P, DNA (plasmid cp32-6 gene
             800927-35-9P, DNA (plasmid cp32-3 gene BBS36 fragment)
     BBM10)
     800927-37-1P, DNA (plasmid cp32-3 gene BBS36) 800927-39-3P
                  800927-43-9P 800927-45-1P
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     800927-41-7P
                                                              800927-49-5P,
     DNA (plasmid lp54 gene BBA04 fragment) 800927-51-9P, DNA (plasmid lp54
     gene BBA04) 800927-53-1P 800927-55-3P 800927-57-5P, DNA (plasmid
     cp26 gene BBB14 fragment) 800927-59-7P, DNA (plasmid cp26 gene BBB14)
                  800927-63-3P 800927-65-5P, DNA (plasmid lp28-1 gene
     800927-61-1P
      ***vls***
                fragment) 800927-67-7P, DNA (plasmid lp28-1 gene vls8
     fragment) 800927-69-9P, DNA (plasmid lp28-1 gene vls9 fragment)
     800927-71-3P
                  800927-72-4P, DNA (plasmid lp28-1 gene vlpE1)
     800927-74-6P
                   800927-76-8P 800927-78-0P 800927-80-4P 800927-81-5P
     800927-83-7P, DNA (plasmid lp5 gene BBT01 fragment) 800927-85-9P, DNA
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(plasmid lp5 gene BBT01) 800927-87-1P 800927-88-2P 800927-90-6P
     800927-92-8P
     RL: BPN (Biosynthetic preparation); DGN (Diagnostic use); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (nucleotide sequence; nucleic acid and/or polypeptide sequences of
          ***Borrelia*** burgdorferi for vaccination and antibody prepn.
        techniques)
ΤТ
     800932-74-5 800932-75-6
     RL: PRP (Properties)
        (unclaimed nucleotide sequence; nucleic acid and/or polypeptide
        sequences of ***Borrelia*** burgdorferi for vaccination and
        antibody prepn. techniques)
L4
     ANSWER 18 OF 87 CAPLUS COPYRIGHT 2009 ACS on STN
ΑN
     2004:565053 CAPLUS <<LOGINID::20090609>>
     141:118336
DN
ΤI
    Polynucleotide and polypeptide sequences for ***vls*** genes of
     pathogenic
                 ***Borrelia***
                                 and their diagnostic and therapeutic uses
     against infection and Lyme disease
IN
     Norris, Steven J.
     Board of Regents, University of Texas System, USA
PA
SO
     PCT Int. Appl., 182 pp.
     CODEN: PIXXD2
DT
    Patent
LA
    English
FAN.CNT 1
     PATENT NO.
                  KIND DATE APPLICATION NO. DATE
                       ____

      WO 2004058181
      A2
      20040715
      WO 2003-US41182
      20031222

      WO 2004058181
      A3
      20050421

PΙ
         W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
             CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
             GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC,
             LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO,
            NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ,
             TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
         RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ,
             BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE,
             ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK,
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    AU 2003299872
                        A1
                               20040722 AU 2003-299872
                                                                 20031222
     EP 1572714
                         Α2
                               20050914 EP 2003-800145
                                                                  20031222
         R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK
                     A1
     US 20060240035
                                20061026
                                         US 2005-539956
                        P
                                20021220
PRAI US 2002-435077P
     WO 2003-US41182
                        W
                                20031222
     The invention claims DNA sequences encoding variable major protein (
AΒ
      ***VMP*** )-like polypeptides of pathogenic ***Borrelia*** , the use
     of the DNA sequences in recombinant vectors to express polypeptides, the
     encoded amino acid sequences, application of the DNA and amino acid
     sequences to the prodn. of polypeptides as antigens for immunoprophylaxis,
     immunotherapy, and immunodiagnosis. The invention also claims use of the
     nucleic acid sequences as probes or primers for the detection of organisms
     causing Lyme disease, relapsing fever, or related disorders, and kits
     designed to facilitate methods of using the described polypeptides, DNA
     segments, and antibodies. Examples of the invention show reactivity of
```

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human Lyme disease serum with recombinant ***Borrelia*** afzelii
      ***Vls*** (variable major protein-like sequence) protein ***VLS***
    -BA13 and with recombinant B. garinii ***Vls*** protein
                                                               ***VLS***
    -BG10. Mouse anti- ***Borrelia*** burgdorferi serum also reacted in an
    enzyme immunoassay with the recombinant proteins ***VLS*** -BA13 and
      ***VLS*** -BG10. The examples also show gene organization of
    silent cassette loci from B. afzelii strain ACAI and B. garinii strain
    Ip90, expression of gene vlsE, and cDNA sequences of vlsE variants cloned
    from strains that were passaged through mice.
RE.CNT 2
             THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD
             ALL CITATIONS AVAILABLE IN THE RE FORMAT
    Polynucleotide and polypeptide sequences for ***vls*** genes of
    pathogenic
               ***Borrelia*** and their diagnostic and therapeutic uses
    against infection and Lyme disease
    The invention claims DNA sequences encoding variable major protein (
      of the DNA sequences in recombinant vectors to express polypeptides, the
    encoded amino acid sequences, application of the. . . the described
    polypeptides, DNA segments, and antibodies. Examples of the invention
    show reactivity of human Lyme disease serum with recombinant
      ***Borrelia*** afzelii ***Vls*** (variable major protein-like
    sequence) protein ***VLS*** -BA13 and with recombinant B. garinii
      ***Vls*** protein ***VLS*** -BG10. Mouse anti- ***Borrelia***
    burgdorferi serum also reacted in an enzyme immunoassay with the
    recombinant proteins ***VLS*** -BA13 and ***VLS*** -BG10. The
    examples also show gene organization of ***vls*** silent cassette loci
    from B. afzelii strain ACAI and B. garinii strain Ip90, expression of gene
    vlsE, and cDNA sequences. .
    DNA sequence ***Borrelia*** gene
                                        ***vls*** antigen;
      ***Borrelia*** gene ***vls*** diagnosis vaccine immunotherapy Lyme
    disease infection
    Infection
       (bacterial; polynucleotide and polypeptide sequences for ***vls***
       genes of pathogenic ***Borrelia*** and their diagnostic and
       therapeutic uses against infection and Lyme disease)
    Immunoassay
       (enzyme-linked immunosorbent assay; polynucleotide and polypeptide
       sequences for ***vls*** genes of pathogenic ***Borrelia*** and
       their diagnostic and therapeutic uses against infection and Lyme
       disease)
    Recombination, genetic
       (gene conversion; polynucleotide and polypeptide sequences for
         ***vls*** genes of pathogenic ***Borrelia*** and their
diagnostic
       and therapeutic uses against infection and Lyme disease)
       (immunodiagnosis; polynucleotide and polypeptide sequences for
         ***vls*** genes of pathogenic ***Borrelia*** and their
       and therapeutic uses against infection and Lyme disease)
    Animals
    Bos taurus
    Canis familiaris
    Cervidae
    Equus caballus
    Human
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ΤI

AB

ST

ΤТ

ΙT

ΤT

ΙT

ΙT

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Mus
        (infection; polynucleotide and polypeptide sequences for ***vls***
       genes of pathogenic ***Borrelia*** and their diagnostic and
       therapeutic uses against infection and Lyme disease)
ΤT
    Antibodies and Immunoglobulins
    RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); DGN
     (Diagnostic use); ANST (Analytical study); BIOL (Biological study); USES
     (Uses)
        (labeled; polynucleotide and polypeptide sequences for
       genes of pathogenic ***Borrelia*** and their diagnostic and
       therapeutic uses against infection and Lyme disease)
    Antibodies and Immunoglobulins
ΙT
     RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); DGN
     (Diagnostic use); ANST (Analytical study); BIOL (Biological study); USES
     (Uses)
        (monoclonal; polynucleotide and polypeptide sequences for
       genes of pathogenic ***Borrelia*** and their diagnostic and
       therapeutic uses against infection and Lyme disease)
ΤТ
    Antigenic variation
     Blood analysis
         ***Borrelia***
                        afzelii
        ***Borrelia*** burgdorferi
         ***Borrelia*** garinii
     DNA sequences
     Genetic polymorphism
     Immunity
     Immunoassay
     Immunoblotting
     Immunoprecipitation
     Immunotherapy
    Lyme disease
    Molecular cloning
    Nucleic acid amplification (method)
     Plasmids
    Protein sequences
    Radioimmunoassay
     Test kits
    Urine analysis
     cDNA sequences
        (polynucleotide and polypeptide sequences for ***vls***
       pathogenic ***Borrelia*** and their diagnostic and therapeutic uses
       against infection and Lyme disease)
ΤТ
    Antigens
    RL: ANT (Analyte); BPN (Biosynthetic preparation); DGN (Diagnostic use);
     PRP (Properties); ANST (Analytical study); BIOL (Biological study); PREP
     (Preparation); USES (Uses)
        (polynucleotide and polypeptide sequences for ***vls***
       pathogenic ***Borrelia*** and their diagnostic and therapeutic uses
       against infection and Lyme disease)
    Nucleic acids
ΤТ
     RL: ANT (Analyte); BUU (Biological use, unclassified); DGN (Diagnostic
     use); ANST (Analytical study); BIOL (Biological study); USES (Uses)
        (polynucleotide and polypeptide sequences for ***vls*** genes of
       pathogenic
                   ***Borrelia***
                                    and their diagnostic and therapeutic uses
       against infection and Lyme disease)
    Antibodies and Immunoglobulins
ΙT
```

```
RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); DGN
     (Diagnostic use); ANST (Analytical study); BIOL (Biological study); USES
        (polynucleotide and polypeptide sequences for ***vls***
                                                                   genes of
                   ***Borrelia***
       pathogenic
                                    and their diagnostic and therapeutic uses
       against infection and Lyme disease)
    Primers (nucleic acid)
    RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); DGN
     (Diagnostic use); PRP (Properties); ANST (Analytical study); BIOL
     (Biological study); USES (Uses)
        (polynucleotide and polypeptide sequences for ***vls*** genes of
       pathogenic
                   ***Borrelia*** and their diagnostic and therapeutic uses
       against infection and Lyme disease)
ΙT
    Escherichia coli
        (recombinant host; polynucleotide and polypeptide sequences for
          ***vls*** genes of pathogenic ***Borrelia*** and their
diagnostic
       and therapeutic uses against infection and Lyme disease)
ΤТ
    Fever and Hyperthermia
        (relapsing; polynucleotide and polypeptide sequences for
       genes of pathogenic ***Borrelia*** and their diagnostic and
       therapeutic uses against infection and Lyme disease)
ΙT
    Repetitive DNA
    RL: BSU (Biological study, unclassified); PRP (Properties); BIOL
     (Biological study)
        ( ***vls***
                     silent cassettes; polynucleotide and polypeptide
       sequences for ***vls*** genes of pathogenic ***Borrelia***
       their diagnostic and therapeutic uses against infection and Lyme
       disease)
ΤТ
    Gene, microbial
     RL: ANT (Analyte); BUU (Biological use, unclassified); DGN (Diagnostic
     use); PRP (Properties); ANST (Analytical study); BIOL (Biological study);
     USES (Uses)
        ( ***vls*** ; polynucleotide and polypeptide sequences for
          ***vls*** genes of pathogenic ***Borrelia***
                                                           and their
diagnostic
       and therapeutic uses against infection and Lyme disease)
ΙT
     Gene, microbial
     RL: ANT (Analyte); BUU (Biological use, unclassified); DGN (Diagnostic
     use); PRP (Properties); ANST (Analytical study); BIOL (Biological study);
     USES (Uses)
        (vlsE; polynucleotide and polypeptide sequences for ***vls*** genes
       of pathogenic ***Borrelia*** and their diagnostic and therapeutic
       uses against infection and Lyme disease)
ΙT
    721865-74-3
                  721865-75-4
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                                              721865-92-5
     RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); DGN
     (Diagnostic use); PRP (Properties); ANST (Analytical study); BIOL
     (Biological study); USES (Uses)
        ( ***Borrelia***
                          afzelii strain ACAI gene vls13 primer;
       polynucleotide and polypeptide sequences for ***vls*** genes of
       pathogenic ***Borrelia*** and their diagnostic and therapeutic uses
       against infection and Lyme disease)
ΤТ
    721865-72-1
    RL: ANT (Analyte); BUU (Biological use, unclassified); DGN (Diagnostic
     use); PRP (Properties); ANST (Analytical study); BIOL (Biological study);
    USES (Uses)
        ( ***Borrelia*** burgdorferi B31 vlsE and ***vls*** silent
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cassette flanking direct repeat; polynucleotide and polypeptide
       sequences for ***vls*** genes of pathogenic ***Borrelia*** and
       their diagnostic and therapeutic uses against infection and Lyme
       disease)
    721865-93-6 721865-94-7 721865-95-8
                                            721865-96-9
ΤT
    RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); DGN
     (Diagnostic use); PRP (Properties); ANST (Analytical study); BIOL
     (Biological study); USES (Uses)
       ( ***Borrelia*** garinii strain Ip90 gene vls10 primer;
       polynucleotide and polypeptide sequences for ***vls*** genes of
       pathogenic ***Borrelia*** and their diagnostic and therapeutic uses
       against infection and Lyme disease)
ΙT
    721863-14-5
    RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); DGN
     (Diagnostic use); PRP (Properties); ANST (Analytical study); BIOL
     (Biological study); USES (Uses)
       ( ***Borrelia*** gene ***vls*** specific primer 4470;
       polynucleotide and polypeptide sequences for ***vls*** genes of
       pathogenic ***Borrelia*** and their diagnostic and therapeutic uses
       against infection and Lyme disease)
    721863-15-6
ΙT
     RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); DGN
     (Diagnostic use); PRP (Properties); ANST (Analytical study); BIOL
     (Biological study); USES (Uses)
       ( ***Borrelia*** gene ***vls*** specific primer 4471;
       polynucleotide and polypeptide sequences for ***vls*** genes of
       pathogenic ***Borrelia*** and their diagnostic and therapeutic uses
       against infection and Lyme disease)
ΙT
    721863-03-2
     RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); DGN
     (Diagnostic use); PRP (Properties); ANST (Analytical study); BIOL
     (Biological study); USES (Uses)
       ( ***Borrelia*** gene ***vls***
                                            specific primer 4540;
       polynucleotide and polypeptide sequences for ***vls*** genes of
       pathogenic ***Borrelia*** and their diagnostic and therapeutic uses
       against infection and Lyme disease)
ΙT
    721863-11-2
    RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); DGN
     (Diagnostic use); PRP (Properties); ANST (Analytical study); BIOL
     (Biological study); USES (Uses)
       ( ***Borrelia*** gene ***vls*** specific primer 4545;
       polynucleotide and polypeptide sequences for ***vls*** genes of
       pathogenic ***Borrelia*** and their diagnostic and therapeutic uses
       against infection and Lyme disease)
ΙT
    721863-10-1
     RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); DGN
     (Diagnostic use); PRP (Properties); ANST (Analytical study); BIOL
     (Biological study); USES (Uses)
       ( ***Borrelia*** gene ***vls*** specific primer 4548;
       polynucleotide and polypeptide sequences for ***vls*** genes of
       pathogenic ***Borrelia*** and their diagnostic and therapeutic uses
       against infection and Lyme disease)
ΤТ
    721863-12-3
     RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); DGN
     (Diagnostic use); PRP (Properties); ANST (Analytical study); BIOL
     (Biological study); USES (Uses)
        ( ***Borrelia*** gene ***vls*** specific primer 4587;
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polynucleotide and polypeptide sequences for ***vls*** genes of
       pathogenic ***Borrelia*** and their diagnostic and therapeutic uses
       against infection and Lyme disease)
ΙT
    721863-13-4
    RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); DGN
     (Diagnostic use); PRP (Properties); ANST (Analytical study); BIOL
     (Biological study); USES (Uses)
       ( ***Borrelia*** gene ***vls*** specific primer 4588;
       polynucleotide and polypeptide sequences for ***vls*** genes of
       pathogenic ***Borrelia*** and their diagnostic and therapeutic uses
       against infection and Lyme disease)
    721862-92-6P 721862-95-9P
                                721863-00-9P
                                               721863-01-0P
                                                              721863-02-1P
ΙT
    721863-07-6P 721863-08-7P 721863-09-8P 721863-19-0P 721863-20-3P
    721863-33-8P 721863-34-9P 721863-35-0P 721863-36-1P 721863-37-2P
    721863-38-3P 721863-39-4P 721863-40-7P 721863-41-8P 721863-42-9P
    721863-43-0P 721863-45-2P 721863-48-5P 721863-62-3P
                                                              721863-63-4P
    721863-64-5P 721863-65-6P 721863-66-7P 721863-67-8P 721863-70-3P 721863-73-6P 721863-74-7P 721865-61-8P, Antigen
                                                721863-67-8P 721863-68-9P
    (plasmid pBG-10-1 gene vls10) 721865-63-0P 721865-64-1P 721865-65-2P
    721865-66-3P 721865-67-4P 721865-68-5P 721865-71-0P, Antigen
    (plasmid pBA-13-1 gene vls13)
    RL: ANT (Analyte); BPN (Biosynthetic preparation); DGN (Diagnostic use);
    PRP (Properties); ANST (Analytical study); BIOL (Biological study); PREP
    (Preparation); USES (Uses)
       (amino acid sequence; polynucleotide and polypeptide sequences for
         ***vls*** genes of pathogenic ***Borrelia*** and their
diagnostic
       and therapeutic uses against infection and Lyme disease)
    511612-64-9 511612-65-0 511612-66-1 511612-67-2 511612-68-3
ΤТ
                              511612-71-8 511612-72-9
                                                         511612-73-0
                 511612-70-7
    511612-69-4
                511612-75-2 511612-76-3 511612-77-4 511612-78-5
    511612-74-1
    511612-79-6, Antigen ( ***Borrelia*** afzelii strain ACAI clone 2622
    gene vlsE C-terminal fragment) 511612-80-9, Antigen ( ***Borrelia***
    afzelii strain ACAI clone 2624a gene vlsE C-terminal fragment)
    511612-81-0, Antigen ( ***Borrelia*** afzelii strain ACAI clone 2624b
    gene vlsE C-terminal fragment) 511612-82-1, Antigen ( ***Borrelia***
    afzelii strain ACAI clone 2625 gene vlsE fragment) 511612-83-2
    511612-84-3 511612-85-4 511612-86-5
                                           511612-87-6 511612-88-7
                                           511612-92-3 511612-93-4
    511612-89-8 511612-90-1 511612-91-2
    511612-94-5 511612-95-6 511612-96-7, Antigen ( ***Borrelia***
    garinii strain Ip90 clone 17 gene vlsE fragment) 511612-97-8, Antigen (
      ***Borrelia*** garinii strain Ip90 clone 20 gene vlsE fragment)
    511612-98-9, Antigen ( ***Borrelia*** garinii strain Ip90 clone 21 gene
    vlsE fragment) 511612-99-0, Antigen ( ***Borrelia*** garinii strain
    Ip90 clone 23 gene vlsE fragment)
    RL: BSU (Biological study, unclassified); PRP (Properties); BIOL
     (Biological study)
       (amino acid sequence; polynucleotide and polypeptide sequences for
         ***vls*** genes of pathogenic ***Borrelia*** and their
diagnostic
       and therapeutic uses against infection and Lyme disease)
ΙT
    721862-91-5 721862-96-0 721862-97-1 721862-98-2 721862-99-3
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    721863-27-0 721863-28-1 721863-29-2 721863-30-5 721863-31-6
    721863-32-7 721863-44-1 721863-46-3 721863-47-4 721863-49-6
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721863-55-4 721863-56-5 721863-57-6 721863-58-7 721863-59-8
     721863-60-1 721863-61-2 721863-69-0 721863-71-4 721863-72-5
     721865-60-7, DNA (plasmid pBG-10-1 gene vls10)
                                                    721865-62-9, DNA (plasmid
    pBA-13-1 gene vls13) 721865-69-6 721865-70-9
    RL: ANT (Analyte); BUU (Biological use, unclassified); DGN (Diagnostic
    use); PRP (Properties); ANST (Analytical study); BIOL (Biological study);
     USES (Uses)
        (nucleotide sequence; polynucleotide and polypeptide sequences for
         ***vls*** genes of pathogenic ***Borrelia*** and their
diagnostic
       and therapeutic uses against infection and Lyme disease)
     503713-49-3 503713-50-6 503713-51-7 503713-52-8 503713-53-9
     503713-54-0 503713-55-1 503713-56-2 503713-57-3 503713-58-4
     RL: BSU (Biological study, unclassified); PRP (Properties); BIOL
     (Biological study)
        (nucleotide sequence; polynucleotide and polypeptide sequences for
         ***vls*** genes of pathogenic ***Borrelia*** and their
diagnostic
       and therapeutic uses against infection and Lyme disease)
     58-85-5D, Biotin, conjugates
     RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); DGN
     (Diagnostic use); ANST (Analytical study); BIOL (Biological study); USES
     (Uses)
       (polynucleotide and polypeptide sequences for ***vls***
       pathogenic ***Borrelia*** and their diagnostic and therapeutic uses
       against infection and Lyme disease)
     145856-09-3, GenBank L04788 391840-97-4, GenBank U76405
     RL: BSU (Biological study, unclassified); PRP (Properties); BIOL
     (Biological study)
        (polynucleotide and polypeptide sequences for ***vls*** genes of
       pathogenic
                   ***Borrelia***
                                    and their diagnostic and therapeutic uses
       against infection and Lyme disease)
    721865-73-2
     RL: BUU (Biological use, unclassified); PRP (Properties); BIOL (Biological
     study); USES (Uses)
       (restriction endonuclease EcoRI-site linker; polynucleotide and
       polypeptide sequences for ***vls*** genes of pathogenic
         ***Borrelia*** and their diagnostic and therapeutic uses against
       infection and Lyme disease)
                 721869-22-3 721869-24-5
    721869-20-1
     RL: PRP (Properties)
        (unclaimed nucleotide sequence; polynucleotide and polypeptide
       sequences for ***vls*** genes of pathogenic ***Borrelia*** and
       their diagnostic and therapeutic uses against infection and Lyme
       disease)
    721869-21-2
                  721869-23-4
     RL: PRP (Properties)
       (unclaimed protein sequence; polynucleotide and polypeptide sequences
       for ***vls*** genes of pathogenic ***Borrelia*** and their
       diagnostic and therapeutic uses against infection and Lyme disease)
     116934-33-9
    RL: PRP (Properties)
       (unclaimed sequence; polynucleotide and polypeptide sequences for
         ***vls*** genes of pathogenic ***Borrelia*** and their
       and therapeutic uses against infection and Lyme disease)
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- L4 ANSWER 19 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN DUPLICATE 11
- AN 2005:50985 BIOSIS <<LOGINID::20090609>>
- DN PREV200500047406
- TI Effects of vlsE complementation on the infectivity of ***Borrelia*** burgdorferi lacking the linear plasmid lp28-1.
- AU Lawrenz, Matthew B.; Wooten, R. Mark; Norris, Steven J. [Reprint Author]
- CS Sch MedDept Pathol and Lab Med, Univ Texas, POB 20708, Houston, TX, 77225, USA
 - steven.j.norris@uth.tmc.edu
- SO Infection and Immunity, (November 2004) Vol. 72, No. 11, pp. 6577-6585. print.

 ISSN: 0019-9567 (ISSN print).
- DT Article
- LA English
- ED Entered STN: 26 Jan 2005 Last Updated on STN: 26 Jan 2005
- AΒ The loss of linear plasmid lp28-1, which contains the ***vls*** antigenic variation locus, is associated with reduced infectivity of ***Borrelia*** burgdorfieri in immunocompetent mice. The recombinant shuttle vector pBBE22, which includes the virulence determinant BBE22 from lp25 and restores infectivity to readily transformable B. burgdorferi lacking 1p25 and 1p56, was used to determine the effect of trans expression of vlsE on virulence. Spirochetes lacking lp28-1 were complemented with the plasmid pBBE22:vlsE, containing both BBE22 and vlsE. VlsE protein produced by this construct was expressed and surface accessible in in vitro-cultured B. burgdorferi, as determined by surface proteolysis and immunoblot analysis. Clones lacking 1p25 but containing 1p28-1 and either pBBE22 or pBBE22:vlsE were reisolated consistently from immunocompetent mice 8 weeks after infection. In contrast, a clone lacking both 1p25 and 1p28-1 and complemented with pBBE22:vlsE was isolated from only a single tissue of one of six C3H/HeN mice 8 weeks postinfection. These results indicate that either an intact v/s antigenic variation locus or another determinant on 1p28-1 is required to restore complete infectivity. In addition, an isogenic clone that retained lp28-1 was complemented with the v/sE shuttle plasmid and was examined for vlsEsequence variation and infectivity. Sequence variation was not observed for the shuttle plasmid, indicating that the cis arrangement of v/sE and the ***vls*** silent cassettes in 1p28-1 facilitate vlsE gene conversion. Lack of vlsE sequence variation on the shuttle plasmid thus did not result in clearance of the trans-complemented strain in immunocompetent mice under the conditions tested.
- TI Effects of vlsE complementation on the infectivity of ***Borrelia*** burgdorferi lacking the linear plasmid lp28-1.
- AB The loss of linear plasmid lp28-1, which contains the ***vls*** antigenic variation locus, is associated with reduced infectivity of ***Borrelia*** burgdorfieri in immunocompetent mice. The recombinant shuttle vector pBBE22, which includes the virulence determinant BBE22 from lp25 and restores infectivity. . . and infectivity. Sequence variation was not observed for the shuttle plasmid, indicating that the cis arrangement of v/sE and the ***vls*** silent cassettes in lp28-1 facilitate vlsE gene conversion. Lack of vlsE sequence variation on the shuttle plasmid thus did not. .

ORGN . . .

Mammals, Nonhuman Vertebrates, Nonhuman Mammals, Rodents, Vertebrates ORGN Classifier

Spirochaetaceae 06112

Super Taxa

Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name

Borrelia burgdorferi (species): pathogen

Taxa Notes

Bacteria, Eubacteria, Microorganisms

- L4 ANSWER 20 OF 87 MEDLINE on STN
- AN 2004243144 MEDLINE <<LOGINID::20090609>>
- DN PubMed ID: 15128807
- TI Lysine-dependent multipoint binding of the ***Borrelia*** burgdorferi virulence factor outer surface protein E to the C terminus of factor H.
- AU Alitalo Antti; Meri Taru; Chen Tong; Lankinen Hilkka; Cheng Zhu-Zhu; Jokiranta T Sakari; Seppala Ilkka J T; Lahdenne Pekka; Hefty P Scott; Akins Darrin R; Meri Seppo
- CS Department of Bacteriology and Immunology, Haartman Institute and Helsinki University Central Hospital, University of Helsinki, Helsinki, Finland.
- NC AI-07364 (United States NIAID NIH HHS) RR-15564 (United States NCRR NIH HHS)
- SO Journal of immunology (Baltimore, Md.: 1950), (2004 May 15) Vol. 172, No. 10, pp. 6195-201.

 Journal code: 2985117R. ISSN: 0022-1767.
- CY United States
- DT Journal; Article; (JOURNAL ARTICLE) (RESEARCH SUPPORT, NON-U.S. GOV'T) (RESEARCH SUPPORT, U.S. GOV'T, P.H.S.)
- LA English
- FS Abridged Index Medicus Journals; Priority Journals
- EM 200409
- ED Entered STN: 15 May 2004
 Last Updated on STN: 3 Sep 2004
 Entered Medline: 2 Sep 2004
- Serum resistance, an important virulence determinant of ***Borrelia*** AB burgdorferi sensu lato strains belonging to the ***Borrelia*** afzelii and B. burgdorferi sensu stricto genotypes, is related to binding of the complement inhibitor factor H to the spirochete surface protein outer surface protein E (OspE) and its homologues. In this study, we show that the C-terminal short consensus repeats 18-20 of both human and mouse factor H bind to OspE. Analogously, factor H-related protein 1, a distinct plasma protein with three short consensus repeat domains homologous to those in factor H, bound to OspE. Deleting 15-aa residues (region V) from the C terminus of the OspE paralog P21 (a 20.7-kDa OspE-paralogous surface lipoprotein in the B. burgdorferi sensu stricto 297 strain) abolished factor H binding. However, C-terminal peptides from OspE, P21, or OspEF-related protein P alone and the C-terminal deletion mutants of P21 inhibited factor H binding to OspE only partially when compared with full-length P21 or its N-terminal mutant. Alanine substitution of amino acids in peptides from the key binding regions of the OspE family indicated that several lysine residues are required for factor H binding. Thus, the ***borrelial*** OspE family proteins bind the C inhibitor factor H via multiple sites in a lysine-dependent manner. The C-terminal site V (Ala(151)-Lys(166)) is necessary, but not sufficient, for factor H binding in both rodents and humans. Identification of the necessary binding sites forms a basis for the development of vaccines that block the factor H-OspE interaction and thereby promote the killing of ***Borreliae*** .
- TI Lysine-dependent multipoint binding of the ***Borrelia*** burgdorferi

virulence factor outer surface protein E to the C terminus of factor H. Serum resistance, an important virulence determinant of ***Borrelia*** AB burgdorferi sensu lato strains belonging to the ***Borrelia*** and B. burgdorferi sensu stricto genotypes, is related to binding of the complement inhibitor factor H to the spirochete. . . key binding regions of the OspE family indicated that several lysine residues are required for factor H binding. Thus, the ***borrelial*** proteins bind the C inhibitor factor H via multiple sites in a lysine-dependent manner. The C-terminal site V. . . forms a basis for the development of vaccines that block the factor H-OspE interaction and thereby promote the killing of ***Borreliae*** CT . . GE, genetics *Bacterial Outer Membrane Proteins: ME, metabolism Bacterial Proteins: GE, genetics *Bacterial Proteins: ME, metabolism Blood Proteins: ME, metabolism *** Borrelia burgdorferi: GE, genetics*** ****Borrelia burgdorferi: ME, metabolism*** *** Borrelia burgdorferi: PY, pathogenicity*** Complement Factor H: AI, antagonists & inhibitors *Complement Factor H: ME, metabolism Consensus Sequence Heparin: PD, . . ***133483-07-5 (VMP21 antigen, Borrelia)*** ; 56-87-1 (Lysine); RN 80295-65-4 (Complement Factor H); 9005-49-6 (Heparin) 0 (Antigens, Bacterial); 0 (Bacterial Outer Membrane Proteins); 0 CN (Bacterial Proteins); 0 (Blood Proteins); 0 (Lipoproteins); 0 (OspE ***Borrelia*** burgdorferi); 0 (Peptide Fragments); 0 protein, (Virulence Factors); 0 (complement factor H, human); 0 (factor H-related protein 1) ANSWER 21 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on L42005:31487 BIOSIS <<LOGINID::20090609>> ΑN DN PREV200500031306 TΙ ***Borrelia*** burgdorferi changes its surface antigenic expression in response to host immune responses. Liang, Fang Ting; Yan, Jun; Mbow, M. Lamine; Sviat, Steven L.; Gilmore, ΑU Robert D.; Mamula, Mark; Fikriq, Erol [Reprint Author] CS Sch MedDept Internal MedRheumatol Sect, Yale Univ, S525A,300 Cedar St, New Haven, CT, 06520, USA erol.fikriq@yale.edu Infection and Immunity, (October 2004) Vol. 72, No. 10, pp. 5759-5767. SO ISSN: 0019-9567 (ISSN print). DТ Article LA English Entered STN: 12 Jan 2005 EDLast Updated on STN: 12 Jan 2005 The Lyme disease spirochete, ***Borrelia*** burgdorferi, causes AΒ persistent mammalian infection despite the development of vigorous immune responses against the pathogen. To examine spirochetal phenotypes that dominate in the hostile immune environment, the mRNA transcripts of four prototypic surface lipoproteins, decorin-binding protein A (DbpA), outer surface protein C (OspC), BBF01, and VlsE, were analyzed by quantitative reverse transcription-PCR under various immune conditions. We demonstrate that B. burgdorferi changes its surface antigenic expression in response

to immune attack. dbpA expression was unchanged while the spirochetes decreased ospC expression by 446 times and increased BBF01 and vlsE expression up to 20 and 32 times, respectively, under the influence of immune pressure generated in inummocompetent mice during infection. This change in antigenic expression could be induced by passively immunizing infected severe combined immunodeficiency mice with specific antisera or OspC antibody and appears to allow B. ***Borrelia*** burgdorferi to resist immune attack. ***Borrelia*** burgdorferi changes its surface antigenic expression in response to host immune responses. The Lyme disease spirochete, ***Borrelia*** burgdorferi, causes persistent mammalian infection despite the development of vigorous immune responses against the pathogen. To examine spirochetal phenotypes that. . . during infection. This change in antigenic expression could be induced by passively immunizing infected severe combined immunodeficiency mice with specific ***Borrelia*** antisera or OspC antibody and appears to allow B. burgdorferi to resist immune attack. Lyme disease: bacterial disease, complications, etiology, immunology, pathology Lyme Disease (MeSH) Chemicals & Biochemicals BBF01 [VlsE]: expression, prototypic surface lipoprotein; ***Vmp*** -like sequence, expressed [VlsE]; decorin-binding protein A [DbpA]: expression, prototypic surface lipoprotein; mRNA [messenger RNA]; outer surface protein C [OspC]: expression,. . . Mammals, Nonhuman Vertebrates, Nonhuman Mammals, Rodents, Vertebrates ORGN Classifier Spirochaetaceae 06112 Super Taxa Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name ***Borrelia*** burgdorferi (species): pathogen, strain-B31 clone 5A11 Taxa Notes Bacteria, Eubacteria, Microorganisms ***Borrelia*** burgdorferi BBF01 gene (Spirochaetaceae); ***Borrelia*** burgdorferi actin gene (Spirochaetaceae); ***Borrelia*** burgdorferi dbpA gene [***Borrelia*** burgdorferi decorin-binding protein A gene] (Spirochaetaceae); ***Borrelia*** burgdorferi flaB gene (Spirochaetaceae); ***Borrelia*** burgdorferi ospC gene [***Borrelia*** burgdorferi outer surface protein A gene] (Spirochaetaceae); ***Borrelia*** burgdorferi vlsE gene [***Borrelia*** burgdorferi ***Vmp*** -like sequence, expressed gene] (Spirochaetaceae) ANSWER 22 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on DUPLICATE 12 2003:153698 BIOSIS <<LOGINID::20090609>> PREV200300153698 Characterization of the ***vls*** antigenic variation loci of the Lyme disease spirochaetes ***Borrelia*** qarinii Ip90 and ***Borrelia***

Wang, Dachun; Botkin, Douglas J.; Norris, Steven J. [Reprint Author]

Houston, University of Texas, PO Box 20708, Houston, TX, 77225-0708, USA

Department of Pathology and Laboratory Medicine, Medical School at

ΤI

AB

ΙT

ΤT

GEN

L4

ΑN DN

ΤI

ΑU

CS

afzelii ACAI.

Steven.J.Norris@uth.tmc.edu

- SO Molecular Microbiology, (March 2003) Vol. 47, No. 5, pp. 1407-1417. print. ISSN: 0950-382X (ISSN print).
- DT Article
- LA English
- ED Entered STN: 26 Mar 2003 Last Updated on STN: 26 Mar 2003
- AB The ***vls*** locus of ***Borrelia*** burgdorferi B31 consists of 15 silent cassettes and one expression site (vlsE), and the presence of the encoding plasmid Ip28-1 correlates with high infectivity. Recombination between the expression cassette and silent cassettes occurs in vivo, and this process may enable B. burgdorferi to evade the immune response. To determine the characteristics of the ***vls*** loci in ***Borrelia*** strains, we have cloned and characterized the ***vls*** silent cassette loci of ***Borrelia*** garinii Ip90 and ***Borrelia*** afzelii ACAI, consisting of 11 ***vls*** silent ***vls*** silent cassettes respectively. The silent cassettes and 14 cassettes share 90-97% nucleotide sequence identity with one another within the Ip90 ***vls*** locus and 84-91% within the ACAI ***vls*** locus. In both organisms, the silent cassettes resemble the B31 ***Vls*** sequences in overall amino acid similarity (50-65%) and in

the

- presence of six variable regions interspersed between six relatively invariant regions. The vise expression sites of these two strains have not been isolated, but transcripts of vise were detected by reverse transcriptase-polymerase chain reaction for both Ip90 and ACAI. In addition, the occurrence of sequence variation within the vlsE cassette region of these transcripts was verified. This study indicates that the ***vls*** loci present in B. garinii Ip90 and B. afzelii ACAI have characteristics similar to those found in B. burgdorferi B31.
- TI Characterization of the ***vls*** antigenic variation loci of the Lyme disease spirochaetes ***Borrelia*** garinii Ip90 and ***Borrelia*** afzelii ACAI.
- AB The ***vls*** locus of ***Borrelia*** burgdorferi B31 consists of 15 silent cassettes and one expression site (vlsE), and the presence of the encoding plasmid Ip28-1. . . in vivo, and this process may enable B. burgdorferi to evade the immune response. To determine the characteristics of the ***vls*** loci in other ***Borrelia*** strains, we have cloned and characterized the ***vls*** silent cassette loci of ***Borrelia*** garinii Ip90 and ***Borrelia*** afzelii ACAI, consisting of 11 ***vls*** silent cassettes and 14 ***vls*** silent cassettes respectively. The silent cassettes share 90-97% nucleotide sequence identity with one another within the Ip90 ***vls*** locus and 84-91% within the ACAI ***vls*** locus. In

both

organisms, the silent cassettes resemble the B31 ***Vls*** sequences in overall amino acid similarity (50-65%) and in the presence of six variable regions interspersed between six relatively invariant. . . the occurrence of sequence variation within the vlsE cassette region of these transcripts was verified. This study indicates that the ***vls*** loci present in B. garinii Ip90 and B. afzelii ACAI have characteristics similar to those found in B. burgdorferi B31.

ORGN Classifier

Spirochaetaceae 06112

Super Taxa

Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name

```
***Borrelia***
***Borrelia***
burgdorferi (species): parasite, strain-ACAI
***Borrelia***
garinii (species): parasite, strain-Ip90
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Taxa Notes

Bacteria, Eubacteria, Microorganisms

GEN ***vls*** gene: antigenic variation loci

- L4 ANSWER 23 OF 87 CABA COPYRIGHT 2009 CABI on STN DUPLICATE 13
- AN 2003:147583 CABA <<LOGINID::20090609>>
- DN 20033120690
- TI Evaluation of a canine C6 ELISA Lyme disease test for the determination of the infection status of cats naturally exposed to ***Borrelia*** burgdorferi
- AU Levy, S. A.; O'Connor, T. E.; Hanscom, J. L.; Shields, P.
- CS Durham Veterinary Hospital PC, 178 Parmelee Hill Road, Durham, CT 06422, USA.
- SO Veterinary Therapeutics, (2003) Vol. 4, No. 2, pp. 172-177. 27 ref. Publisher: Veterinary Learning Systems Inc. Trenton ISSN: 1528-3593
- CY United States
- DT Journal
- LA English
- ED Entered STN: 16 Sep 2003 Last Updated on STN: 16 Sep 2003
- The efficacy of a commercially available in-office kit (SNAP 3Dx, IDEXX AB Laboratories) for detection of antibodies directed against an invariable region (IR6) of the B. burgdorferi surface protein VlsE (***Vmp*** -like sequence, Ex pressed), a surface antigen of the spirochete recognized during active infection has been evaluated in dogs. The present study was conducted to determine whether this in-office test could be useful for detection of antibodies to B. burgdorferi in cats. Cats owned by clients of a veterinary hospital located in an area hyperendemic for Lyme disease were included in the study. When possible, cats with an outdoor lifestyle, bite wounds, or current tick infestation were recruited for the study to help ensure that animals with a likelihood of exposure to natural infection by B. burgdorferi would be included in the test group. Of the 24 cats tested, 17 samples were positive for antibodies to B. burgdorferi by the C6 ELISA kit. For all 17 of these samples, a duplicate sample tested by immunofluorescent assay (IFA) was in agreement with the ELISA. Five samples were negative by both assays. Two samples that were negative by the C6 ELISA test had low IFA titers (1:100). One of these two discrepant samples was negative and one was positive for antibodies to B. burgdorferi by the Western blot test. It was concluded that the C6 ELISA test performed with good agreement with the IFA and Western blot tests for detection of antibody to B. burgdorferi in the majority of cats tested. The test offers the advantages of producing a result rapidly (approximately 8 minutes), and it requires only two drops of serum, plasma, or whole blood.
- TI . . . of a canine C6 ELISA Lyme disease test for the determination of the infection status of cats naturally exposed to $**Borrelia*** burgdorferi.$
- AB . . . 3Dx, IDEXX Laboratories) for detection of antibodies directed against an invariable region (IR6) of the B. burgdorferi surface protein VlsE (***Vmp*** -like sequence, Ex pressed), a surface antigen of the spirochete recognized during active infection has been evaluated in dogs. The present. . .
- BT ***Borrelia*** ; Spirochaetaceae; Spirochaetales; Gracilicutes;

bacteria; prokaryotes; Felis; Felidae; Fissipeda; carnivores; mammals; vertebrates; Chordata; animals; small mammals ORGN ***Borrelia*** burgdorferi; cats ANSWER 24 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on

- T.4
- ΑN 2002:523404 BIOSIS <<LOGINID::20090609>>
- DNPREV200200523404
- TΤ ***VMP*** -like sequences of pathogenic ***borrelia***
- ΑU Norris, Steven J. [Inventor, Reprint author]; Zhang, Jing-Ren [Inventor]; Hardham, John M. [Inventor]; Howell, Jerrilyn K. [Inventor]; Barbour, Alan G. [Inventor]; Weinstock, George M. [Inventor]
- CS Houston, TX, USA
 - ASSIGNEE: Board of Regents, The University of Texas System
- PΤ US 6437116 20020820
- Official Gazette of the United States Patent and Trademark Office Patents, SO (Aug. 20, 2002) Vol. 1261, No. 3. http://www.uspto.gov/web/menu/patdata.html. e-file. CODEN: OGUPE7. ISSN: 0098-1133.
- DT Patent
- LA English
- Entered STN: 9 Oct 2002 ED Last Updated on STN: 9 Oct 2002
- The present invention relates to DNA sequences encoding ***Vmp*** -like AΒ polypeptides of pathogenic ***Borrelia*** , the use of the DNA sequences in recombinant vectors to express polypeptides, the encoded amino acid sequences, application of the DNA and amino acid sequences to the production of polypeptides as antigens for immunoprophylaxis, immunotherapy, and immunodiagnosis. Also disclosed are the use of the nucleic acid sequences as probes or primers for the deletion of organisms causing Lyme disease, relapsing fever, or related disorders, and kits designed to facilitate methods of using the described polypeptides, DNA segments and antibodies.
- ΤI ***VMP*** -like sequences of pathogenic ***borrelia*** .
- The present invention relates to DNA sequences encoding ***Vmp*** -like AΒ polypeptides of pathogenic ***Borrelia*** , the use of the DNA sequences in recombinant vectors to express polypeptides, the encoded amino acid sequences, application of the.
- ΤТ Major Concepts

Infection; Molecular Genetics (Biochemistry and Molecular Biophysics)

Chemicals & Biochemicals

DNA sequences; ***Vmp*** -like polypeptides

ORGN Classifier

Spirochaetaceae 06112

Super Taxa

Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name

Borrelia : pathogen

Taxa Notes

Bacteria, Eubacteria, Microorganisms

- ANSWER 25 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on L4DUPLICATE 14
- ΑN 2002:452514 BIOSIS <<LOGINID::20090609>>
- PREV200200452514
- TΤ Evidence that the variable regions of the central domain of VlsE are antigenic during infection with Lyme disease spirochetes.

- AU McDowell, John V.; Sung, Shian-Ying; Hu, Linden T.; Marconi, Richard T. [Reprint author]
- CS Department of Microbiology and Immunology, Medical College of Virginia at Virginia Commonwealth University, Richmond, VA, 23298-0678, USA rmarconi@hsc.vcu.edu
- SO Infection and Immunity, (August, 2002) Vol. 70, No. 8, pp. 4196-4203. print.

 CODEN: INFIBR. ISSN: 0019-9567.
- DT Article
- LA English
- ED Entered STN: 21 Aug 2002 Last Updated on STN: 21 Aug 2002
- AΒ It has been postulated that the ***vls*** system of the Lyme disease spirochetes contributes to immune evasion through antigenic variation. While it is clear that vlsE undergoes sequence change within its variable regions at a high frequency during the early stages of infection, a definitive role in immune evasion has not been demonstrated. In this report we assessed the murine and human humoral immune response to recombinant (r)-VlsE variants that originally arose during infection in mice. Immunoblot analyses of r-VlsE variants were conducted by using serum samples collected from mice infected with ***Borrelia*** burgdorferi clones that carried different vlsE variants. All of the r-VlsE variants were recognized by infection sera regardless of the identity of the infecting clone or isolate. In addition, all variants were immunoreactive with a panel of human Lyme disease patient serum samples. It is evident from these analyses that the infection-induced VlsE variants share common epitopes that reside within conserved segments of these proteins. However, preabsorption experiments revealed that the variable regions of the central domain of VlsE, which undergo rapid mutation during infection, also influence the antigenic properties of the protein. A subset of the antibodies elicited against vlsE variants that differ in the sequences of their variable regions were found to be variant specific. Hence, in spite of a robust antibody response to conserved segments of VlsE, infection-induced sequence changes within the variable regions alter the antigenicity of VlsE. These results provide the first direct evidence of antigenic variation in the VlsE protein.
- AB It has been postulated that the ***vls*** system of the Lyme disease spirochetes contributes to immune evasion through antigenic variation. While it is clear that vlsE undergoes. . . during infection in mice. Immunoblot analyses of r-VlsE variants were conducted by using serum samples collected from mice infected with ***Borrelia*** burgdorferi clones that carried different vlsE variants. All of the r-VlsE variants were recognized by infection sera regardless of the. . .
- GEN ***Borrelia*** burgdorferis vlsE gene (Spirochaetaceae)
- L4 ANSWER 26 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN DUPLICATE 15
- AN 2002:231270 BIOSIS <<LOGINID::20090609>>
- DN PREV200200231270
- TI The 44-kb linear plasmid molecule in the relapsing fever agent ***Borrelia*** duttonii strain Ly serve as a preservation of ***vmp***

genes.

- AU Tabuchi, Norihiko; Mitani, Harumi; Seino, Satoshi; Fukunaga, Masahito [Reprint author]
- CS Laboratory of Molecular Microbiology, Faculty of Pharmacy and Pharmaceutical Sciences, Fukuyama University, Gakuencho 1, Fukuyama,

Hiroshima, 729-0292, Japan mfukunaq@superniq.niq.ac.jp SO Microbiology and Immunology, (2002) Vol. 46, No. 3, pp. 159-165. print. CODEN: MIIMDV. ISSN: 0385-5600. DТ Article LA English ED Entered STN: 3 Apr 2002 Last Updated on STN: 3 Apr 2002 ***Borrelia*** duttonii strain Ly, a causative agent of relapsing AΒ fever, contains a linear one megabase chromosome and 12 linear plasmid molecules. Here we report that the sequence of the 44-kb linear plasmid of strain Ly is found to contain variable major protein (***vmp***) genes for antigenic variation of relapsing fever ***borreliae*** . The determined sequence is of 44,010 bp except for both ends of the molecule. Of 39 open reading frames (ORFs) found in the sequence, 21 ORFs (named ***vmpA*** to U) showed moderate similarities with ***vmp*** genes ***Borrelia*** hermsii. However, most of the ***vmp*** homologues are apparently nonfunctional because of their frameshifts within the sequence and/or absence of promoter and ribosome-binding signals upstream of their genes. RT-PCR experiments using the specific ***vmp*** gene revealed that ***vmpE*** , one of primer for each the ***vmp*** genes, was expressed at the location of the 44-kb plasmid molecule. The result suggests that the plasmid molecule may play a role in the preservation of the serotype switching of ***vmp*** genes in a mammalian host. The 44-kb linear plasmid molecule in the relapsing fever agent ТΤ ***Borrelia*** duttonii strain Ly serve as a preservation of ***amv*** ***Borrelia*** duttonii strain Ly, a causative agent of relapsing AΒ fever, contains a linear one megabase chromosome and 12 linear plasmid molecules.. . . we report that the sequence of the 44-kb linear plasmid of strain Ly is found to contain variable major protein (***vmp***) genes for antigenic variation of relapsing fever ***borreliae*** . The determined sequence is of 44,010 bp except for both ends of the molecule. Of 39 open reading frames (ORFs) found in the sequence, 21 ORFs (named ***vmpA*** to U) showed moderate similarities with ***vmp*** genes ***Borrelia*** hermsii. However, most of the ***vmp*** homologues are apparently nonfunctional because of their frameshifts within the sequence and/or absence of promoter and ribosome-binding signals upstream of their genes. RT-PCR experiments using the specific primer for each ***vmp*** gene revealed that ***vmpE*** , one of the ***vmp*** genes, was expressed at the location of the 44-kb plasmid molecule. The result suggests that the plasmid molecule may play a role in the preservation of the serotype switching of ***vmp*** genes in a mammalian host. ITRT-PCR [reverse transcriptase-polymerase chain reaction]: analytical method, genetic method, polymerase chain reaction ΙT Miscellaneous Descriptors open reading frames; plasmid sequence; ***vmp*** gene preservation [variable major protein gene preservation] ORGN . Notes Animals, Chordates, Humans, Mammals, Primates, Vertebrates ORGN Classifier Spirochaetaceae 06112

Super Taxa

Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name

Borrelia duttonii: pathogen, strain-Ly

Taxa Notes

- L4 ANSWER 27 OF 87 CAPLUS COPYRIGHT 2009 ACS on STN
- AN 2001:791027 CAPLUS <<LOGINID::20090609>>
- DN 136:304895
- TI Analysis of ***Borrelia*** burgdorferi vlsE gene expression and recombination in the tick vector
- AU Indest, Karl J.; Howell, Jerrilyn K.; Jacobs, Mary B.; Scholl-Meeker, Dorothy; Norris, Steven J.; Philipp, Mario T.
- CS Department of Parasitology, Tulane Regional Primate Research Center, Tulane University Health Sciences Center, Covington, LA, 70433, USA
- SO Infection and Immunity (2001), 69(11), 7083-7090 CODEN: INFIBR; ISSN: 0019-9567
- PB American Society for Microbiology
- DT Journal
- LA English
- AΒ Expression and recombination of the antigenic variation vlsE gene of the Lyme disease spirochete ***Borrelia*** burgdorferi were analyzed in the tick vector. To assess vlsE expression, Ixodes scapularis nymphs infected with the B. burgdorferi strain B31 were fed on mice for 48 or 96 h or to repletion, and then crushed and acetone fixed either immediately thereafter (ticks collected at the two earlier time points) or 4 days after repletion. Unfed nymphs also were examd. At all of the time points investigated, spirochetes were able to bind a rabbit antibody raised against the conserved invariable region 6 of VlsE, as assessed by indirect immunofluorescence, but not pre-immune serum from the same rabbit. This same antibody also bound to B31 spirochetes cultivated in vitro. Intensity of fluorescence appeared highest in cultured spirochetes, followed by spirochetes present in unfed ticks. Only a dim fluorescent signal was obsd. on spirochetes at the 48 and 96 h time points and at day 4 post-repletion. Expression of vlsE in vitro was affected by a rise in pH from 7.0 to 8.0 at 34.degree.C. Hence, vlsE expression appears to be sensitive to environmental cues of the type found in the B. burgdorferi natural history. To assess vlsE recombination, nymphs were capillary fed the B. burgdorferi B31 clonal isolate 5A3. Ticks thus infected were either left to rest for 4 wk (Group I) or fed to repletion on a mouse (Group II). The contents of each tick from both groups were cultured and 10 B. burgdorferi clones from the spirochetal isolate of each tick were obtained. The vlsE cassettes from several of these clones were amplified by PCR and sequenced. Regardless of whether the isolate was derived from Group I or Group II ticks, no changes were obsd. in the vlsE sequence. In contrast, vlsE cassettes amplified from B. burgdorferi clones derived from a mouse that was infected with B31-5A3 capillary-fed nymphs showed considerable recombination. It follows that vlsE recombination does not occur in the tick vector.
- RE.CNT 33 THERE ARE 33 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- TI Analysis of ***Borrelia*** burgdorferi vlsE gene expression and recombination in the tick vector
- AB Expression and recombination of the antigenic variation vlsE gene of the Lyme disease spirochete ***Borrelia*** burgdorferi were analyzed in the tick vector. To assess vlsE expression, Ixodes scapularis nymphs

infected with the B. burgdorferi strain. . . DNA sequence ***Borrelia*** gene vlsE mouse infection tick; ST ***Borrelia*** gene vlsE lipoprotein tick expression; protein sequence gene vlsE lipoprotein ***Borrelia*** ; genetic recombination ***Borrelia*** gene vlsE mouse infection tick ΙT Ixodes scapularis (anal. of ***Borrelia*** burgdorferi vlsE gene expression and recombination in tick vector) ΙT Lipoproteins RL: BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study) ***Vmp*** -like sequence; partial sequence and (gene vlsE, for expression in tick of ***Borrelia*** burgdorferi gene vlsE lipoprotein) ΤТ Development, nonmammalian postembryonic (nymph; anal. of ***Borrelia*** burgdorferi vlsE gene expression and recombination in tick vector) ΙT ***Borrelia*** burgdorferi DNA sequences Protein sequences (partial sequence of ***Borrelia*** burgdorferi gene vlsE lipoprotein isolated from mouse infected by infestation with Ixodes scapularis nymphal ticks) ΤТ Lyme disease Mus Recombination, genetic (vlsE cassettes amplified from ***Borrelia*** burgdorferi clones derived from mouse infected with B31-5A3 capillary-fed nymphs showed considerable recombination) ΙT Gene, microbial RL: BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study) (vlsE; partial DNA sequence, expression and recombination in tick vector of ***Borrelia*** burgdorferi gene vlsE) 411243-31-7 411243-32-8 411243-33-9 411243-34-0 ΙT RL: BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study) (amino acid sequence; partial sequence of ***Borrelia*** burgdorferi gene vlsE lipoprotein isolated from mouse infected by infestation with Ixodes scapularis nymphal ticks) 359572-33-1, GenBank AY043397 359572-34-2, GenBank AY043398 359572-35-3, GenBank AY043399 359572-36-4, GenBank AY043400 382261-30-5, GenBank AY043401 RL: BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study) (nucleotide sequence; partial DNA sequence, expression and recombination in tick vector of ***Borrelia*** burgdorferi gene vlsE) ANSWER 28 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on L4DUPLICATE 16 2001:494794 BIOSIS <<LOGINID::20090609>> ΑN DN PREV200100494794 ΤI Evidence for the contribution of point mutations to vlsE variation and for apparent constraints on the net accumulation of sequence changes in vlsE during infection with Lyme disease spirochetes.

Sung, Shian-Ying; McDowell, John V.; Marconi, Richard T. [Reprint author]

ΑU

- CS Department of Microbiology and Immunology, Medical College of Virginia at Virginia Commonwealth University, Richmond, VA, 23298-0678, USA rmarconi@hsc.vcu.edu
- SO Journal of Bacteriology, (October, 2001) Vol. 183, No. 20, pp. 5855-5861. print. CODEN: JOBAAY. ISSN: 0021-9193.
- DT Article
- LA English
- OS Genbank-AF354775; Genbank-AF354776; Genbank-AF354777; Genbank-AF354778; Genbank-AF354779; Genbank-AF354780; Genbank-AF354781; Genbank-AF354782; Genbank-AF354783; Genbank-AF354784; Genbank-AF354785; Genbank-AF354786; Genbank-AF354787; Genbank-AF354788; Genbank-AF354789; Genbank-AF354790; Genbank-AF354791; Genbank-AF354792; Genbank-AF354793
- ED Entered STN: 24 Oct 2001 Last Updated on STN: 25 Feb 2002
- In the Lyme disease spirochetes, both the ospE and vlsE gene families have AΒ been demonstrated to undergo sequence variation during infection. To further investigate the mechanisms associated with the generation of ***vls*** variation, single-nucleotide polymorphism and subsequent DNA sequence analyses were performed on the vlsE gene and its paralog, BBJ51, a related gene with a frameshift mutation. These analyses focused on a series of postinfection clonal populations obtained from mice infected ***Borrelia*** burgdorferi B31MIpc or its clonal derivative, B31MIc53. vlsE, but not BBJ51, was found to undergo sequence changes during infection. Consistent with that reported previously (J.-R. Zhang et al., Cell 89:275-285, 1997) many of the sequence changes appear to have arisen through gene conversion events and to be localized to the variable regions of vlsE. However, analysis of the vlsE nucleotide sequences revealed that some sequence changes were the result of point mutations, as these changes did not have potential contributing sources in the ***vls*** cassettes. To determine if sequence changes accumulate in vlsE over long-term infection, the vlsE genes of clonal populations recovered after 7 months of infection in mice were analyzed. While new sequence changes developed, a significant number of these changes resulted in the restoration of the vlsE sequence of the original infecting clone. In addition, we noted that some positions within the variable regions (VR) are stable even though the cassettes possess residues that could contribute to sequence variation through gene conversion. These analyses suggest that the total number of amino acid sequence changes that can be maintained by VlsE levels off during infection. In summary, in this report we demonstrate that the development of point mutations serves as a second mechanism by which vlsE sequence variation can be generated and that the capacity for vlsE variation, while still significant, is less than previously postulated.
- AB. . . families have been demonstrated to undergo sequence variation during infection. To further investigate the mechanisms associated with the ***vls*** variation, single-nucleotide polymorphism and generation of subsequent DNA sequence analyses were performed on the vlsE gene and its paralog, BBJ51, a related. . . gene with a frameshift mutation. These analyses focused on a series of postinfection clonal populations obtained from mice infected with ***Borrelia*** burgdorferi B31MIpc or its clonal derivative, B31MIc53. vlsE, but not BBJ51, was found to undergo sequence changes during infection. Consistent. . . some sequence changes were the result of point mutations, as these changes did not have potential contributing sources in the ***vls*** cassettes. To determine if sequence changes accumulate in vlsE over long-term infection, the vlsE genes of clonal populations recovered after. . .

ORGN . . . Mammals, Nonhuman Vertebrates, Nonhuman Mammals, Rodents, Vertebrates
ORGN Classifier
Spirochaetaceae 06112

Super Taxa

Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name

Borrelia burgdorferi: pathogen

Taxa Notes

Bacteria, Eubacteria, Microorganisms

GEN ***Borrelia*** burgdorferi vlsE gene (Spirochaetaceae): frameshift mutation, point mutation, sequence changes

L4 ANSWER 29 OF 87 CAPLUS COPYRIGHT 2009 ACS on STN

AN 2001:695183 CAPLUS <<LOGINID::20090609>>

DN 135:370510

- TI ***Borrelia*** burgdorferi-induced inflammation facilitates spirochete adaptation and variable major protein-like sequence locus recombination
- AU Anguita, Juan; Thomas, Venetta; Samanta, Swapna; Persinski, Rafal; Hernanz, Carmen; Barthold, Stephen W.; Fikrig, Erol
- CS Section of Rheumatology, Department of Internal Medicine, Yale University School of Medicine, New Haven, CT, 06520, USA
- SO Journal of Immunology (2001), 167(6), 3383-3390 CODEN: JOIMA3; ISSN: 0022-1767
- PB American Association of Immunologists
- DT Journal
- LA English
- AB Spirochete adaptation in vivo is assocd. with preferential B. burgdorferi gene expression. Here, the authors show that the administration of B. burgdorferi-immune sera to IFN-.gamma.R-deficient mice that have been infected with B. burgdorferi N40 for 4 days causes spirochete clearance. In contrast, immune sera-mediated clearance of B. burgdorferi N40 is not apparent in immunocompetent mice, suggesting a role for IFN-.gamma.-mediated responses in B. burgdorferi N40 host adaptation. B. burgdorferi-immune sera also induce clearance of B. burgdorferi N40 that have been passaged in vitro 75 times (B. burgdorferi N40-75), a deriv. of B. burgdorferi N40 that does not rapidly adapt in vivo in immunocompetent mice. B. burgdorferi N40-75 produces lower levels of IFN-.gamma. and IL-12 in mice than does B. burgdorferi N40, and the administration of these cytokines to B. burgdorferi N40-75-infected mice results in an increased spirochetal burden, further indicating that IFN-.gamma.-mediated events promote B. burgdorferi survival. Differential immunoscreening and RT-PCR demonstrate that IFN-.gamma.-mediated signals facilitate spirochete recombination at the variable major protein like sequence locus, a site for early antigenic variation in vivo, and that recombination rates by B. burgdorferi N40 are lower in IFN-.gamma.R-deficient mice than in control animals. Thus, the murine immune response can promote the in vivo adaptation of B. burgdorferi.
- RE.CNT 60 THERE ARE 60 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- TI ***Borrelia*** burgdorferi-induced inflammation facilitates spirochete adaptation and variable major protein-like sequence locus recombination
- ST ***Borrelia*** adaptation Lyme disease ***vls*** gene recombination cytokine
- IT Adaptation, microbial

Borrelia burgdorferi

Inflammation

Recombination, genetic

(***Borrelia*** burgdorferi-induced inflammation facilitates
spirochete adaptation and variable major protein-like sequence locus
recombination)

IT Signal transduction, biological

IT Interleukin 12

RL: BAC (Biological activity or effector, except adverse); BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)

(interferon .gamma./interleukin-12 signals facilitate ***Borrelia*** burgdorferi recombination at variable major protein like sequence locus)

IT Gene, microbial

RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)

IT Interferons

RL: BAC (Biological activity or effector, except adverse); BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)

(.gamma.; interferon .gamma. signals facilitate ***Borrelia*** burgdorferi recombination at variable major protein like sequence locus)

- L4 ANSWER 30 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN
- AN 2002:222937 BIOSIS <<LOGINID::20090609>>
- DN PREV200200222937
- TI Comparative analysis of the kinetics of mutation in the ***vls*** and ospE antigenic variation systems of Lyme disease spirochetes during infection in mice.
- AU Sung, S. [Reprint author]; McDowell, J. V. [Reprint author]; Marconi, R. T. [Reprint author]
- CS Medical College of VA of VCU, Richmond, VA, USA
- SO Abstracts of the General Meeting of the American Society for Microbiology, (2001) Vol. 101, pp. 325-326. print.

Meeting Info.: 101st General Meeting of the American Society for Microbiology. Orlando, FL, USA. May 20-24, 2001. American Society of Microbiology.

ISSN: 1060-2011.

- DT Conference; (Meeting)
 Conference; Abstract; (Meeting Abstract)
- LA English
- ED Entered STN: 3 Apr 2002 Last Updated on STN: 3 Apr 2002

although the majority of the genome sequence of B31MI has been determined, the segment thought to carry vlsE was neither sequenced or mapped. PCR analyses confirmed the presence of a vlsE and demonstrated that this allele was maintained after 3 months of infection in mice. PCR and hybridization analyses demonstrated that large scale rearrangements in vlsE did not occur over the course of infection. Hybridization analyses of DNA fractionated by two dimensional pulsed field gel electrophoresis demonstrated vlsE to be carried on the linear plasmid, 1p28-1. To assess vlsE genetic stability, vlsE was amplified from post-infection clones recovered from ear punch biopsies from mice infected with B31MI and the amplicons were screened for polymorphisms. Genetic changes were detected and a comparision of the mutation rates in vlsE and ospE revealed that the vlsE gene undergoes mutation at a higher frequency than ospE. A silent B. burgdorferi B31MI ***vls*** allele, vlsE1, which harbors an frameshift mutation, was found to be genetically stable. To determine if mutations continue to accumulate over long term infection, a clone recovered from the ear punch biopsy was used to infect a mouse and the infection was allowed to persist for 7 months. Analysis of vlsE from the clonal populations recovered at 7 months revealed that while some new mutations developed many of the genetic changes were in the form of reversions. These analyses suggest that there is a limit to the accumulation of mutations in the vlsE gene and that the potential number of variants that can arise may not be as great as previously postulated.

- TI Comparative analysis of the kinetics of mutation in the ***vls*** and ospE antigenic variation systems of Lyme disease spirochetes during infection in mice.

ORGN . .

Mammals, Nonhuman Vertebrates, Nonhuman Mammals, Rodents, Vertebrates ORGN Classifier

Spirochaetaceae 06112

Super Taxa

Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name

Borrelia burgdorferi: pathogen

Taxa Notes

- GEN ***Borrelia*** burgdorferi ospE gene (Spirochaetaceae): analysis, mutation; ***Borrelia*** burgdorferi vlsE gene (Spirochaetaceae): analysis, mutation
- L4 ANSWER 31 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN DUPLICATE 17
- AN 2001:372097 BIOSIS <<LOGINID::20090609>>
- DN PREV200100372097
- TI Increased expression of ***Borrelia*** burgdorferi vlsE in response to human endothelial cell membranes.
- AU Hudson, Charlene R.; Frye, Jonathan G.; Quinn, Frederick D.; Gherardini, Frank C. [Reprint author]
- CS Department of Microbiology, University of Georgia, 546 Biological Sciences

Building, Athens, GA, 30602, USA FRANKG@arches.uga.edu

- SO Molecular Microbiology, (July, 2001) Vol. 41, No. 1, pp. 229-239. print. CODEN: MOMIEE. ISSN: 0950-382X.
- DT Article
- LA English
- OS Genbank-AF314755
- ED Entered STN: 8 Aug 2001 Last Updated on STN: 19 Feb 2002
- AΒ RNA isolated from virulent ***Borrelia*** burgdorferi cells incubated with human endothelial or neurological tissue cells was subjected to subtractive hybridization using RNA from the same strain incubated in tissue culture medium alone. This RNA subtractive technique generated specific probes that hybridized to two restriction fragments (8.2 kb and 10 kb respectively) generated by EcoRI digestion of total plasmid DNA. The 10 kb EcoRI fragment localized to Ip28-1 and was subsequently identified as the variable membrane protein-like sequence (***vls***) region, which includes an expression locus (vlsE) and 15 silent cassettes. vlsE encodes a 36 kDa outer surface protein that undergoes antigenic variation during animal infections. Primer extension analysis identified the 5' end of a transcript and a putative promoter for vlsE. Quantitative reverse transcription-polymerase chain reaction (RT-PCR) suggested that the expression of vlsE increased when virulent B. burgdorferi cells were incubated with human tissue cells or purified cell membranes isolated from those same cell lines. A 138 bp region upstream of the vlsE region that was not reported in the genome sequence was sequenced using specific 32P end-labelled primers in a DNA cycle sequencing system at high annealing temperatures. Analysis revealed that it contained a 51 bp inverted repeat, which could form an extremely stable cruciform structure. Southern blots probed with the vlsE promoter/operator region indicated that part or all of this sequence could be found on other B. burgdorferi plasmids.
- TI Increased expression of ***Borrelia*** burgdorferi vlsE in response to human endothelial cell membranes.
- AB RNA isolated from virulent ***Borrelia*** burgdorferi cells incubated with human endothelial or neurological tissue cells was subjected to subtractive hybridization using RNA from the same. . . plasmid DNA. The 10 kb EcoRI fragment localized to Ip28-1 and was subsequently identified as the variable membrane protein-like sequence (***vls***) region, which includes an expression locus (vlsE) and 15 silent cassettes. vlsE encodes a 36 kDa outer surface protein that. . .
- GEN ***Borrelia*** burgdorferi vlsE gene (Spirochaetaceae): expression
- L4 ANSWER 32 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN DUPLICATE 18
- AN 2001:355831 BIOSIS <<LOGINID::20090609>>
- DN PREV200100355831
- TI Analysis of a ***VMP*** -like sequence (***vls***) locus in
 Borrelia garinii and ***Vls*** homologues among four
 Borrelia burgdorferi sensu lato species.
- AU Wang, Guiqing [Reprint author]; van Dam, Alje P.; Dankert, Jacob
- CS Department of Biochemistry and Molecular Biology, New York Medical College, Valhalla, NY, 10595, USA guiqing_wang@nymc.edu
- SO FEMS Microbiology Letters, (15 May, 2001) Vol. 199, No. 1, pp. 39-45. print.
 - CODEN: FMLED7. ISSN: 0378-1097.

- DT Article
- LA English
- OS Genbank-AF274070; Genbank-AF274071; Genbank-AF274072; Genbank-AF274073; Genbank-AF274074; Genbank-AF274075; Genbank-AF274076
- ED Entered STN: 2 Aug 2001 Last Updated on STN: 23 Feb 2002
- AB ***VMP*** -like sequence (***vls***) locus that consists of one expressed vlsE gene and 15 silent ***vls*** cassettes has been described in ***Borrelia*** burgdorferi sensu stricto B31. In the present study, the ***vls*** locus from a ***Borrelia*** isolate A87SA was analyzed. DNA fragments that contained three complete and five partial ***vls*** cassettes were cloned and sequenced. Pulsed-field gel electrophoresis (PFGE) analysis and Southern hybridization of the PFGE blot indicated that the B. garinii A87SA, consisting of at least eight ***vls*** cassettes, was located on a 21-kb linear plasmid. The size of the three complete ***vls*** cassettes varied from 573 to 612 bp. They had 93.8-94.3% identity at the nucleotide level and 84.9-87.3% amino acid identity. The amino acid sequences of the three ***vls*** cassettes of B. garinii A87SA exhibited 45.9-50.8% identity to the VlsE sequence of B. burgdorferi B31, and 30.0-33.8% identity to the ***VMP17*** sequence of B. hermsii HS1. Homologues of the ***vls*** locus of B. garinii were detected by dot blot hybridization among 24 of the 30 (80.0%) isolates representing four B. burgdorferi sensu lato species distributed widely in Europe. Our findings indicate that B. garinii might possess a similar ***vls*** structure to that described in B. burgdorferi sensu stricto. The highly conserved nature of the ***vls*** locus among various B. burgdorferi sensu lato species suggests that it may be important in the physiology and pathogenesis of Lyme disease spirochetes.
- TI Analysis of a ***VMP*** -like sequence (***vls***) locus in
 Borrelia garinii and ***Vls*** homologues among four
 Borrelia burgdorferi sensu lato species.
- ***VMP*** -like sequence (***vls***) locus that consists of one AΒ expressed vlsE gene and 15 silent ***vls*** cassettes has been described in ***Borrelia*** burgdorferi sensu stricto B31. In the present study, the ***vls*** locus from a ***Borrelia*** garinii isolate A87SA was analyzed. DNA fragments that contained three complete and five partial ***vls*** cassettes were cloned and sequenced. Pulsed-field gel electrophoresis (PFGE) analysis and Southern hybridization of the PFGE blot indicated that the ***vls*** B. garinii A87SA, consisting of at least eight ***vls*** cassettes, was located on a 21-kb linear plasmid. The size of the three complete ***vls*** cassettes varied from 573 to 612 bp. They had 93.8-94.3% identity at the nucleotide level and 84.9-87.3% amino acid identity. The amino acid sequences of the three ***vls*** cassettes of B. garinii A87SA exhibited 45.9-50.8% identity to the VlsE sequence of B. burgdorferi B31, and 30.0-33.8% identity to the ***VMP17*** sequence of B. hermsii HS1. Homologues of the ***vls*** locus of B. garinii were detected by dot blot hybridization among 24 of the 30 (80.0%) isolates representing four B. burgdorferi sensu lato species distributed widely in Europe. Our findings indicate that B. garinii might possess a similar ***vls*** structure to that described in B. burgdorferi sensu stricto. The highly conserved nature of the ***vls*** locus among various B. burgdorferi sensu lato species suggests that it may be important in the physiology and pathogenesis of. . .

ORGN Classifier

```
Super Taxa
       Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms
     Organism Name
           ***Borrelia***
                           burgdorferi: pathogen
           ***Borrelia*** garinii: pathogen
     Taxa Notes
       Bacteria, Eubacteria, Microorganisms
GEN
       ***Borrelia*** garinii ***vls*** gene (Spirochaetaceae);
       ***Borrelia***
                       garinii vlsE gene (Spirochaetaceae)
    ANSWER 33 OF 87 CAPLUS COPYRIGHT 2009 ACS on STN
L4
ΑN
    DN
    134:70362
ΤI
    Combined decorin binding protein and outer surface protein compositions
    and methods of use
    Hanson, Mark S.; Patel, Nita K.; Cassatt, David R.
ΙN
PΑ
    Medimmune, Inc., USA
SO
    PCT Int. Appl., 111 pp.
    CODEN: PIXXD2
DT
    Patent
LA
    English
FAN.CNT 1
                     KIND DATE APPLICATION NO.
    PATENT NO.
                                                                DATE
                       ____
                                           _____

      WO 2000078800
      A2
      20001228
      WO 2000-US16763

      WO 2000078800
      A3
      20010719

                                                                20000616
PΤ
        W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR,
            CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU,
            ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU,
            LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD,
            SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU,
            ZA, ZW
        RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
            DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ,
            CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
PRAI US 1999-140258P P 19990618
    Disclosed are surprisingly effective compns., therapeutic kits and
    vaccines comprising one or more ***Borrelia*** decorin binding protein
     components and one or more ***Borrelia*** outer surface protein
     components. Methods and medical uses are also disclosed in which the
     compns., kits and vaccines are administered to prevent and/or treat
      ***Borrelial*** infections, notably the ***Borrelial*** infections
     that cause Lyme disease.
RE.CNT 3
             THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD
             ALL CITATIONS AVAILABLE IN THE RE FORMAT
AB
    Disclosed are surprisingly effective compns., therapeutic kits and
     vaccines comprising one or more ***Borrelia*** decorin binding protein
     components and one or more ***Borrelia*** outer surface protein
     components. Methods and medical uses are also disclosed in which the
     compns., kits and vaccines are administered to prevent and/or treat
      ***Borrelial*** infections, notably the ***Borrelial*** infections
    that cause Lyme disease.
      ***Borrelia*** decorin binding protein DbpA vaccine; outer surface
ST
    protein OspA ***Borrelia*** vaccine
ΤТ
    Decorins
    RL: BPN (Biosynthetic preparation); THU (Therapeutic use); BIOL
     (Biological study); PREP (Preparation); USES (Uses)
```

(-binding protein; combination of decorin binding protein and outer surface protein as vaccine for preventing or treating ***Borrelia*** infection or Lyme disease) Proteins, specific or class RL: BPN (Biosynthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (DbpA; combination of decorin binding protein and outer surface protein as vaccine for preventing or treating ***Borrelia*** infection or Lyme disease) Proteins, specific or class RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (DbpB; combination of decorin binding protein and outer surface protein as vaccine for preventing or treating ***Borrelia*** infection or Lyme disease) Proteins, specific or class RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (EppA; combination of decorin binding protein and outer surface protein as vaccine for preventing or treating ***Borrelia*** infection or Lyme disease) ***Borrelia*** garinii (G25; combination of decorin binding protein and outer surface protein as vaccine for preventing or treating ***Borrelia*** infection or Lyme disease) Lipoproteins RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (Osp or outer surface proteins; combination of decorin binding protein and outer surface protein as vaccine for preventing or treating ***Borrelia*** infection or Lyme disease) Lipoproteins RL: BPN (Biosynthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (OspA or outer surface protein A; combination of decorin binding protein and outer surface protein as vaccine for preventing or treating ***Borrelia*** infection or Lyme disease) Lipoproteins RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (OspB or outer surface protein B; combination of decorin binding protein and outer surface protein as vaccine for preventing or treating ***Borrelia*** infection or Lyme disease) Lipoproteins RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (OspC or outer surface protein C; combination of decorin binding protein and outer surface protein as vaccine for preventing or treating ***Borrelia*** infection or Lyme disease) Lipoproteins RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (OspD or outer surface protein D; combination of decorin binding protein and outer surface protein as vaccine for preventing or treating ***Borrelia*** infection or Lyme disease) Lipoproteins RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (OspE or outer surface protein E; combination of decorin binding protein and outer surface protein as vaccine for preventing or treating ***Borrelia*** infection or Lyme disease) Lipoproteins RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(OspF or outer surface protein F; combination of decorin binding

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protein and outer surface protein as vaccine for preventing or treating ***Borrelia*** infection or Lyme disease) Proteins, specific or class ΙT RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (S1; combination of decorin binding protein and outer surface protein as vaccine for preventing or treating ***Borrelia*** infection or Lyme disease) ΤT Proteins, specific or class RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (T5; combination of decorin binding protein and outer surface protein as vaccine for preventing or treating ***Borrelia*** infection or Lyme disease) ΙT Proteins, specific or class RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (***Vmp7**** ; combination of decorin binding protein and outer surface protein as vaccine for preventing or treating ***Borrelia*** infection or Lyme disease) ΙT Immunostimulants (adjuvants; combination of decorin binding protein and outer surface protein as vaccine for preventing or treating ***Borrelia*** infection or Lyme disease) ΤТ Animal ***Borrelia*** ***Borrelia*** afzelii ***Borrelia*** burgdorferi Drug delivery systems Lyme disease Molecular cloning Vaccines (combination of decorin binding protein and outer surface protein as vaccine for preventing or treating ***Borrelia*** infection or Lyme disease) Antibodies ΤT RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (combination of decorin binding protein and outer surface protein as vaccine for preventing or treating ***Borrelia*** infection or Lyme disease) ΙT Flagellins RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (combination of decorin binding protein and outer surface protein as vaccine for preventing or treating ***Borrelia*** infection or Lyme disease) ΙT Medical goods (containers; combination of decorin binding protein and outer surface protein as vaccine for preventing or treating ***Borrelia*** infection or Lyme disease) Proteins, specific or class RL: BPN (Biosynthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (decorin-binding; combination of decorin binding protein and outer surface protein as vaccine for preventing or treating ***Borrelia*** infection or Lyme disease) ΙT Immunoglobulins RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (fragments; combination of decorin binding protein and outer surface

protein as vaccine for preventing or treating ***Borrelia***
infection or Lyme disease)

IT Drug delivery systems

(injections, intradermal; combination of decorin binding protein and
outer surface protein as vaccine for preventing or treating
 Borrelia infection or Lyme disease)

IT Drug delivery systems

(injections, s.c.; combination of decorin binding protein and outer surface protein as vaccine for preventing or treating ***Borrelia*** infection or Lyme disease)

IT Containers

(medical; combination of decorin binding protein and outer surface
protein as vaccine for preventing or treating ***Borrelia***
infection or Lyme disease)

IT Drug delivery systems

(nasal, intra-; combination of decorin binding protein and outer surface protein as vaccine for preventing or treating ***Borrelia*** infection or Lyme disease)

IT Proteins, specific or class

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (p110; combination of decorin binding protein and outer surface protein as vaccine for preventing or treating ***Borrelia*** infection or Lyme disease)

IT Proteins, specific or class

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (p13; combination of decorin binding protein and outer surface protein as vaccine for preventing or treating ***Borrelia*** infection or Lyme disease)

IT Proteins, specific or class

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (p17; combination of decorin binding protein and outer surface protein as vaccine for preventing or treating ***Borrelia*** infection or Lyme disease)

IT Proteins, specific or class

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (p28; combination of decorin binding protein and outer surface protein as vaccine for preventing or treating ***Borrelia*** infection or Lyme disease)

IT Proteins, specific or class

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (p35; combination of decorin binding protein and outer surface protein as vaccine for preventing or treating ***Borrelia*** infection or Lyme disease)

IT Proteins, specific or class

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (p37; combination of decorin binding protein and outer surface protein as vaccine for preventing or treating ***Borrelia*** infection or Lyme disease)

IT Proteins, specific or class

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (p39 .alpha.; combination of decorin binding protein and outer surface protein as vaccine for preventing or treating ***Borrelia*** infection or Lyme disease)

IT Proteins, specific or class

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (p39 .beta.; combination of decorin binding protein and outer surface protein as vaccine for preventing or treating ***Borrelia***

infection or Lyme disease)

- IT 7429-90-5, Aluminum, biological studies 21645-51-2, Aluminum hydroxide, biological studies
 - RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (adjuvant; combination of decorin binding protein and outer surface protein as vaccine for preventing or treating ***Borrelia*** infection or Lyme disease)
- L4 ANSWER 34 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN DUPLICATE 19
- AN 2001:63090 BIOSIS <<LOGINID::20090609>>
- DN PREV200100063090
- TI Correlation between plasmid content and infectivity in ***Borrelia*** burgdorferi.
- AU Purser, Joye E.; Norris, Steven J. [Reprint author]
- CS Department of Pathology and Laboratory Medicine, Medical School, and Graduate School of Biomedical, University of Texas-Houston Health Science Center, Houston, TX, 77225, USA Steven.J.Norris@uth.tmc.edu
- SO Proceedings of the National Academy of Sciences of the United States of America, (December 5, 2000) Vol. 97, No. 25, pp. 13865-13870. print. CODEN: PNASA6. ISSN: 0027-8424.
- DT Article
- LA English
- ED Entered STN: 31 Jan 2001 Last Updated on STN: 12 Feb 2002
- AB Infectivity-associated plasmids were identified in ***Borrelia*** burgdorferi B31 by using PCR to detect each of the plasmids in a panel of 19 clonal isolates. The clones exhibited high-, low-, and intermediate-infectivity phenotypes based on their frequency of isolation from needle-inoculated C3H/HeN mice. Presence or absence of 21 of the 22 $\,$ plasmids was determined in each of the clones by using PCR primers specific for regions unique to each plasmid, as identified in the recently available genome sequence. Southern blot hybridization results were used to confirm the PCR results in some cases. Plasmid 1p25 exhibited a direct correlation with infectivity in that it was consistently present in all clones of high or intermediate infectivity and was absent in all low-infectivity clones. lp28-1, containing the ***vmp*** -like sequence locus, also correlated with infectivity; all clones that lacked lp28-1 but contained 1p25 had an intermediate infectivity phenotype, in which infection was primarily restricted to the joints. Plasmids cp9, cp32-3, 1p21, 1p28-2, 1p28-4, and 1p56 apparently are not required for infection in this model, because clones lacking these plasmids exhibited a high-infectivity phenotype. Plasmids cp26, cp32-1, cp32-2 and/or cp32-7, cp32-4, cp32-6, cp32-8, cp32-9, lp17, lp28-3, lp36, lp38, and lp54 were consistently present in all clones examined. On the basis of these results, 1p25 and 1p28-1 appear to encode virulence factors important in the pathogenesis of B. burgdorferi B31.
- TI Correlation between plasmid content and infectivity in ***Borrelia*** burgdorferi.
- AB Infectivity-associated plasmids were identified in ***Borrelia*** burgdorferi B31 by using PCR to detect each of the plasmids in a panel of 19 clonal isolates. The clones. . . consistently present in all clones of high or intermediate infectivity and was absent in all low-infectivity clones. 1p28-1, containing the ***vmp*** -like sequence locus, also correlated with infectivity; all clones that lacked 1p28-1 but contained 1p25 had an intermediate infectivity phenotype, in. . .

ORGN . . . Mammals, Nonhuman Vertebrates, Nonhuman Mammals, Rodents, Vertebrates ORGN Classifier

Spirochaetaceae 06112

Super Taxa

Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name

Borrelia burgdorferi: pathogen

Taxa Notes

Bacteria, Eubacteria, Microorganisms

- L4ANSWER 35 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on DUPLICATE 20
- ΑN 2000:452609 BIOSIS <<LOGINID::20090609>>
- DMPREV200000452609
- Direct evidence for involvement of NF-kappaB in transcriptional activation ТΤ of tumor necrosis factor by a spirochetal lipoprotein.
- ΑU Udalova, Irina A. [Reprint author]; Vidal, Vincent; Scragg, Ian G.; Kwiatkowski, Dominic
- CS Molecular Infectious Disease Group, Institute of Molecular Medicine, John Radcliffe Hospital, Oxford, OX3 9DS, UK
- SO Infection and Immunity, (September, 2000) Vol. 68, No. 9, pp. 5447-5449. print. CODEN: INFIBR. ISSN: 0019-9567.
- DT Article
- T.A English
- ED Entered STN: 25 Oct 2000 Last Updated on STN: 10 Jan 2002
- Variable major lipoprotein (***Vmp***) is a major tumor necrosis AΒ factor (TNF)-inducing component of ***Borrelia*** recurrentis, the agent of louse-borne relapsing fever. B. recurrentis ***Vmp*** rapidly stimulates nuclear translocation of NF-kappaB and proinflammatory cytokine gene expression in the human monocyte-like cell line MonoMac 6. By overexpressing disabled mutant IkappaBalpha in MonoMac 6 cells cotransfected with a reporter gene, we provide evidence that NF-kappaB is essential for the transcriptional activation of TNF in this system.
- Variable major lipoprotein (***Vmp***) is a major tumor necrosis AΒ factor (TNF)-inducing component of ***Borrelia*** recurrentis, the
 agent of louse-borne relapsing fever. B. recurrentis ***Vmp*** rapidly stimulates nuclear translocation of NF-kappaB and proinflammatory cytokine gene expression in the human monocyte-like cell line MonoMac 6. Ву. . .

ORGN . . .

Notes

Animals, Chordates, Humans, Mammals, Primates, Vertebrates ORGN Classifier

Spirochaetaceae 06112

Super Taxa

Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name

Borrelia recurrentis: pathogen

Taxa Notes

Bacteria, Eubacteria, Microorganisms

T.4 ANSWER 36 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN DUPLICATE 21

2000:169153 BIOSIS <<LOGINID::20090609>> ΑN

- DN PREV200000169153
- TI Conservation and heterogeneity of vlsE among human and tick isolates of ***Borrelia*** burgdorferi.
- AU Iyer, Radha; Hardham, John M.; Wormser, Gary P.; Schwartz, Ira; Norris, Steven J. [Reprint author]
- CS Department of Pathology and Laboratory Medicine, University of Texas Medical School at Houston, Houston, TX, 77225, USA
- SO Infection and Immunity, (March, 2000) Vol. 68, No. 3, pp. 1714-1718. print.

 CODEN: INFIBR. ISSN: 0019-9567.
- DT Article
- LA English
- ED Entered STN: 3 May 2000 Last Updated on STN: 4 Jan 2002
- AΒ ***vls*** (variable major protein (***VMP***)-like sequence) locus of ***Borrelia*** burgdorferi encodes an antigenic variation system that closely resembles the ***VMP*** system of relapsing fever ***borreliae*** . To determine whether ***vls*** sequences are present consistently in low-passage, infectious isolates of B. burgdorferi, 22 blood and erythema migrans biopsy isolates from Lyme disease patients in Westchester County, New York, were examined by Southern blot and PCR analysis. Each of the strains contained a single plasmid varying in size from 21 to 38 kb that hybridized strongly with a vlsE probe based on the B. burgdorferi B31 sequence. In contrast, PCR products were obtained with only 10 of the 22 strains when primers corresponding to the 5' and 3' regions of the B31 vlsE sequence outside the variable cassette region were used. Only 2 of 16 B. burgdorferi-infected tick specimens yielded detectable PCR product. Eight of 10 strains that yielded a PCR product under these conditions were type 1 (a genotype with a high rate of dissemination), according to PCR-restriction fragment length polymorphism analysis of intergenic rDNA sequences, whereas the isolates that did not yield vlsE PCR products were either type 2 or type 3. Comparison of the sequences of cloned PCR products from the patient isolates indicated a high degree of identity to the B31 sequence, with most of the differences restricted to the hypervariable regions known to undergo sequence variation. Taken together, these results both reinforce previous evidence that ***vls*** sequences are present consistently in low-passage Lyme disease spirochetes and indicate that both highly conserved and heterogeneous subgroups exist with regard to vlsE sequences.
- TI Conservation and heterogeneity of vlsE among human and tick isolates of ***Borrelia*** burgdorferi.
- (variable major protein (***VMP***)-like sequence) ***vls*** AB The ***Borrelia*** burgdorferi encodes an antigenic variation locus of system that closely resembles the ***VMP*** system of relapsing fever ***borreliae*** . To determine whether ***vls*** sequences are present consistently in low-passage, infectious isolates of B. burgdorferi, 22 blood and erythema migrans biopsy isolates from Lyme. . differences restricted to the hypervariable regions known to undergo sequence variation. Taken together, these results both reinforce previous evidence that ***vls*** sequences are present consistently in low-passage Lyme disease spirochetes and indicate that both highly conserved and heterogeneous subgroups exist with. . .
- IT Major Concepts

Infection

IT Diseases

Lyme disease: bacterial disease

Lyme Disease (MeSH) ΙT Chemicals & Biochemicals ***Borrelia*** burgdorferi ***vls*** gene; ***Borrelia***

burgdorferi vlsE gene

ORGN . Notes

Animals, Chordates, Humans, Mammals, Primates, Vertebrates ORGN Classifier

Spirochaetaceae 06112

Super Taxa

Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name

Borrelia burgdorferi: pathogen

Taxa Notes

- L4ANSWER 37 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on DUPLICATE 22
- 2000:369051 BIOSIS <<LOGINID::20090609>> ΑN
- DN PREV200000369051
- TIStructural characterization of the inflammatory moiety of a variable major lipoprotein of ***Borrelia*** recurrentis.
- Scragg, Ian G.; Kwiatkowski, Dominic [Reprint author]; Vidal, Vincent; ΑU Reason, Andrew; Paxton, Thanai; Panico, Maria; Dell, Ann; Morris, Howard
- CS Dept. of Paediatrics, University of Oxford, John Radcliffe Hospital, Oxford, OX3 9DU, UK
- SO Journal of Biological Chemistry, (January 14, 2000) Vol. 275, No. 2, pp. 937-941. print. CODEN: JBCHA3. ISSN: 0021-9258.
- DT Article
- English LA
- Genbank-AJ237608; EMBL-AJ237608; DDBJ-AJ237608 OS
- Entered STN: 30 Aug 2000 ED Last Updated on STN: 8 Jan 2002
- AΒ Louse-borne relapsing fever, caused by ***Borrelia*** recurrentis, provides one of the best documented examples of the causative role of tumor necrosis factor (TNF) in the pathology of severe infection in humans. We have identified the principal TNF-inducing factor of B. recurrentis as a variable major lipoprotein (***Vmp***). Here we report the complete gene sequence of ***Vmp*** , including its lipoprotein leader sequence. Using metabolically labeled forms of the ***Vmp*** we confirm that the TNF inducing properties are associated with the lipid portion of the molecule. Quadrupole orthogonal time of flight mass spectrometry unequivocally locates the lipidic moiety at the NH2-terminal cysteine of the native polypeptide, and indicates the existence of three forms which are consistent with the structures C16:0, C16:0, C16:0 glyceryl cysteine; C18:1, C16:0, C16:0 glyceryl cysteine; and C18:0, C16:0, C16:0 glyceryl cysteine. These data provide the first direct evidence that the TNF inducing lipid modification of native ***Borrelia*** lipoproteins is a structural homologue of the murein lipoprotein of Escherichia coli.
- ΤI Structural characterization of the inflammatory moiety of a variable major lipoprotein of ***Borrelia*** recurrentis.
- AΒ Louse-borne relapsing fever, caused by ***Borrelia*** recurrentis, provides one of the best documented examples of the causative role of tumor necrosis factor (TNF) in the pathology. . . of severe infection in humans. We have identified the principal TNF-inducing factor of B.

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recurrentis as a variable major lipoprotein ( ***Vmp*** ). Here we
    report the complete gene sequence of ***Vmp*** , including its
    lipoprotein leader sequence. Using metabolically labeled forms of the
                        we confirm that the TNF inducing properties are
    native
             ***qmV***
     associated with the lipid portion of the molecule. Quadrupole orthogonal
     time of. . C18:0, C16:0, C16:0 glyceryl cysteine. These data provide
     the first direct evidence that the TNF inducing lipid modification of
     native ***Borrelia*** lipoproteins is a structural homologue of the
    murein lipoprotein of Escherichia coli.
       and Techniques
    Diseases
       louse borne relapsing fever: bacterial disease
    Chemicals & Biochemicals
       tumor necrosis factor; variable major lipoprotein [ ***Vmp*** ]:
       biochemical structure, structural inflammatory moiety characterization
ORGN Classifier
       Spirochaetaceae
                        06112
     Super Taxa
       Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms
     Organism Name
           ***Borrelia*** recurrentis
     Taxa Notes
       Bacteria, Eubacteria, Microorganisms
    ANSWER 38 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on
    2000:388142 BIOSIS <<LOGINID::20090609>>
    PREV200000388142
    Characteristics of the ***vls*** locus of ***Borrelia*** garinii
    Wang, D. [Reprint author]; Norris, S. J. [Reprint author]
    Univ. of Texas Med. School, Houston, TX, USA
    Abstracts of the General Meeting of the American Society for Microbiology,
    (2000) Vol. 100, pp. 275. print.
    Meeting Info.: 100th General Meeting of the American Society for
    Microbiology. Los Angeles, California, USA. May 21-25, 2000. American
    Society for Microbiology.
    ISSN: 1060-2011.
    Conference; (Meeting)
    Conference; Abstract; (Meeting Abstract)
    English
    Entered STN: 13 Sep 2000
    Last Updated on STN: 8 Jan 2002
    Characteristics of the ***vls*** locus of ***Borrelia*** garinii
    Ip90.
    Miscellaneous Descriptors
       gene expression: analysis; genetic loci: analysis, characterization;
         ***vls*** locus: analysis, characterization; Meeting Abstract
ORGN Classifier
       Spirochaetaceae 06112
     Super Taxa
       Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms
     Organism Name
           ***Borrelia*** garinii: pathogen, strain-Ip90
            ***Borrelia*** spp.: pathogen
     Taxa Notes
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- L4 ANSWER 39 OF 87 CAPLUS COPYRIGHT 2009 ACS on STN DUPLICATE 23
- AN 1999:809495 CAPLUS <<LOGINID::20090609>>
- DN 132:277841
- TI Human antibody responses to VlsE antigenic variation protein of ***Borrelia*** burgdorferi
- AU Lawrenz, M. B.; Hardham, J. M.; Owens, R. T.; Nowakowski, J.; Steere, A. C.; Wormser, G. P.; Norris, S. J.
- CS Departments of Pathology and Laboratory Medicine and Microbiology and Molecular Genetics, University of Texas Medical School at Houston, Houston, TX, 77030, USA
- SO Journal of Clinical Microbiology (1999), 37(12), 3997-4004 CODEN: JCMIDW; ISSN: 0095-1137
- PB American Society for Microbiology
- DT Journal
- LA English
- AΒ VlsE is a 35-kDa surface-exposed lipoprotein of B. burgdorferi that was shown previously to undergo antigenic variation through segmental ***vls*** cassettes with vlsE during exptl. recombination of silent mouse infections. Previous data had indicated that sera from North American Lyme disease patients and exptl. infected animals contained antibodies reactive with VlsE. Here, sera from patients with Lyme disease, syphilis, and autoimmune conditions as well as from healthy controls were examd. for reactivity with VlsE by Western blotting and ELISA. Strong Western blot reactivity to a recombinant VlsE cassette region protein was obtained consistently with Lyme disease sera. Although sera from Lyme disease patients also reacted with a band corresponding to VlsE in B. burgdorferi B31-5A3, interpretation was complicated by low levels of VlsE expression in in vitro-cultured B. burgdorferi and by the presence of comigrating bands. An ELISA using recombinant VlsE was compared with an ELISA using sonically disrupted B. burgdorferi as the antigen. For a total of 93 Lyme disease patient sera examd., the VlsE ELISA yielded sensitivities of 63% for culture-confirmed erythema migrans cases and 92% for later stages, as compared to 61 and 98%, resp., for the "whole-cell" ELISA. The specificities of the two assays with healthy blood donor sera were comparable, but the VlsE ELISA was 90% specific with sera from syphilis patients, compared to 20% specificity for the whole-cell ELISA with this group. Neither assay showed reactivity with a panel of sera from 20 non-Lyme disease arthritis patients or 20 systemic lupus erythematosus patients. Thus, VlsE may be useful in the immunodiagnosis of Lyme disease and may offer greater specificity than ELISAs using whole B. burgdorferi as the antigen.
- RE.CNT 26 THERE ARE 26 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- TI Human antibody responses to VlsE antigenic variation protein of ***Borrelia*** burgdorferi
- AB . . . a 35-kDa surface-exposed lipoprotein of B. burgdorferi that was shown previously to undergo antigenic variation through segmental recombination of silent ***vls*** cassettes with vlsE during exptl. mouse infections. Previous data had indicated that sera from North American Lyme disease patients and. . .
- ST antibody VlsE protein ***Borrelia*** Lyme disease serodiagnosis
- IT Gene, microbial
 - RL: BSU (Biological study, unclassified); BIOL (Biological study)
 (VlsE; Western blot and ELISA for detection of human antibody responses to VlsE antigenic variation protein of ***Borrelia*** burgdorferi)

Antigenic variation ΙT

Blood analysis

Borrelia burgdorferi

Lyme disease

(Western blot and ELISA for detection of human antibody responses to VlsE antigenic variation protein of ***Borrelia*** burgdorferi)

ΤТ Antibodies

RL: ANT (Analyte); ANST (Analytical study)

(Western blot and ELISA for detection of human antibody responses to VlsE antigenic variation protein of ***Borrelia*** burgdorferi)

ΙT Proteins, specific or class

> RL: BSU (Biological study, unclassified); BIOL (Biological study) (gene VlsE; Western blot and ELISA for detection of human antibody responses to VlsE antigenic variation protein of ***Borrelia*** burgdorferi)

ΙT Diagnosis

> (serodiagnosis; Western blot and ELISA for detection of human antibody responses to VlsE antigenic variation protein of ***Borrelia*** burgdorferi)

- ANSWER 40 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on L4DUPLICATE 24
- 1999:265523 BIOSIS <<LOGINID::20090609>> ΑN
- PREV199900265523
- ΤI Specific antibodies reactive with the 22-kilodalton major outer surface protein of ***Borrelia*** anserina Ni-NL protect chicks from infection.
- Sambri, Vittorio; Marangoni, Antonella; Olmo, Andrea; Storni, Elisa; ΑU Montagnani, Marco; Fabbi, Massimo; Cevenini, Roberto [Reprint author]
- Section of Microbiology, DMCSS, University of Bologna, St. Orsola Hospital, via Massarenti 9, 40138, Bologna, Italy
- Infection and Immunity, (May, 1999) Vol. 67, No. 5, pp. 2633-2637. print. SO CODEN: INFIBR. ISSN: 0019-9567.
- DT Article
- LA English
- Entered STN: 15 Jul 1999 ED
 - Last Updated on STN: 15 Jul 1999
- An outer surface lipoprotein or 22 kDa was identified in the avian AΒ pathogen ***Borrelia*** anserina Ni-NL by using antibody preparations reactive with bacterial surface-exposed proteins. Amino acid sequence analysis of the 22-kDa protein demonstrated 90% identity with ***VmpA*** of B. turicatae, suggesting that the protein belongs to the family of 20-kDa outer surface proteins of the genus ***Borrelia*** . All of the 60 chicks intramuscularly treated with antibodies specifically reacting with the 22-kDa protein and infected with strain Ni-NL were completely protected from infection, since no spirochetemia was detected, and from death. Control chicks were treated with immune sera raised against apathogenic strain B. anserina Es, which expresses a prominent 20-kDa polypeptide that is also a member of the ***Vmp*** family but does not cross-react immunologically with the 22-kDa protein of the Ni-NL strain. These animals, infected with B. anserina Ni-NL, showed a high degree of spirochetemia 10 days after infection, and all died between 14 and 21 days after infection. The results showed that the 22-kDa surface protein of B. anserina Ni-NL is a determinant of the pathogenic potential of the strain and also confirmed that only strain-specific antibodies are protective against B. anserina infection.
- Specific antibodies reactive with the 22-kilodalton major outer surface TΙ

AB An outer surface lipoprotein or 22 kDa was identified in the avian pathogen ***Borrelia*** anserina Ni-NL by using antibody preparations reactive with bacterial surface-exposed proteins. Amino acid sequence analysis of the 22-kDa protein demonstrated 90% identity with ***VmpA*** of B. turicatae, suggesting that the protein belongs to the family of 20-kDa outer surface proteins of the genus ***Borrelia***. All of the 60 chicks intramuscularly treated with antibodies specifically reacting with the 22-kDa protein and infected with strain Ni-NL. . . raised against apathogenic strain B. anserina Es, which expresses a prominent 20-kDa polypeptide that is also a member of the ***Vmp*** family but does not cross-react immunologically with the 22-kDa protein of the Ni-NL strain. These animals, infected with B. anserina. . .

IT Major Concepts

Infection

IT Diseases

borrelia infection: bacterial disease
Borrelia Infections (MeSH)

IT Chemicals & Biochemicals

outer surface lipoprotein; ***VmpA***

ORGN . .

Notes

Animals, Birds, Chordates, Nonhuman Vertebrates, Vertebrates ORGN Classifier

Spirochaetaceae 06112

Super Taxa

Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name

> ***Borrelia*** anserina ***Borrelia*** turicatae

Taxa Notes

- L4 ANSWER 41 OF 87 LIFESCI COPYRIGHT 2009 CSA on STN
- AN 1999:65969 LIFESCI <<LOGINID::20090609>>
- TI Specific antibodies reactive with the 22-kilodalton major outer surface protein of ***Borrelia*** anserina Ni-NL protect chicks from infection
- AU Sambri, V.; Marangoni, A.; Olmo, A.; Storni, E.; Montagnani, M.; Fabbi, M.; Cevenini, R.*
- CS Section of Microbiology, DMCSS, University of Bologna, St. Orsola Hospital, via Massarenti 9, 40138 Bologna, Italy; E-mail: cevenini@med.unibo.it
- SO Infection and Immunity [Infect. Immun.], (19990500) vol. 65, no. 5, pp. 2633-2637.
 ISSN: 0019-9567.
- DT Journal
- FS 3
- LA English
- SL English
- AB An outer surface lipoprotein of 22 kDa was identified in the avian pathogen ***Borrelia*** anserina Ni-NL by using antibody preparations reactive with bacterial surface-exposed proteins. Amino acid sequence analysis of the 22-kDa protein demonstrated 90% identity with ***VmpA*** of B. turicatae, suggesting that the protein belongs to the family of 20-kDa outer surface proteins of the genus ***Borrelia*** . All of the 60 chicks intramuscularly treated with antibodies specifically reacting

with the 22-kDa protein and infected with strain Ni-NL were completely protected from infection, since no spirochetemia was detected, and from death. Control chicks were treated with immune sera raised against apathogenic strain B. anserina Es, which expresses a prominent 20-kDa polypeptide that is also a member of the ***Vmp*** family but does not cross-react immunologically with the 22-kDa protein of the Ni-NL strain. These animals, infected with B. anserina Ni-NL, showed a high degree of spirochetemia 10 days after infection, and all died between 14 and 21 days after infection. The results showed that the 22-kDa surface protein of B. anserina Ni-NL is a determinant of the pathogenic potential of the strain and also confirmed that only strain-specific antibodies are protective against B. anserina infection.

- TI Specific antibodies reactive with the 22-kilodalton major outer surface protein of ***Borrelia*** anserina Ni-NL protect chicks from infection
- AB An outer surface lipoprotein of 22 kDa was identified in the avian pathogen ***Borrelia*** anserina Ni-NL by using antibody preparations reactive with bacterial surface-exposed proteins. Amino acid sequence analysis of the 22-kDa protein demonstrated 90% identity with ***VmpA*** of B. turicatae, suggesting that the protein belongs to the family of 20-kDa outer surface proteins of the genus ***Borrelia*** . All of the 60 chicks intramuscularly treated with antibodies specifically reacting with the 22-kDa protein and infected with strain Ni-NL. . . raised against apathogenic strain B. anserina Es, which expresses a prominent 20-kDa polypeptide that is also a member of the ***Vmp*** family but does not cross-react immunologically with the 22-kDa protein of the Ni-NL strain. These animals, infected with B. anserina. .
- L4 ANSWER 42 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN DUPLICATE 25
- AN 1999:342464 BIOSIS <<LOGINID::20090609>>
- DN PREV199900342464
- TI The extended promoters for two outer membrane lipoprotein genes of ***Borrelia*** spp. uniquely include a T-rich region.
- AU Sohaskey, Charles D.; Zuckert, Wolfram R.; Barbour, Alan G. [Reprint author]
- CS Departments of Microbiology and Molecular Genetics and Medicine, University of California Irvine, B240 Med Sci I, Irvine, CA, 92697 4025, USA
- SO Molecular Microbiology, (July, 1999) Vol. 33, No. 1, pp. 41-51. print. CODEN: MOMIEE. ISSN: 0950-382X.
- DT Article
- LA English
- ED Entered STN: 24 Aug 1999 Last Updated on STN: 24 Aug 1999
- AB OspA and B proteins of ***Borrelia*** burgdorferi and ***Vmp*** proteins of ***Borrelia*** hermsii are abundant outer membrane lipoproteins, whose expression varies with the environment. The genes for these proteins have the '-35' and '-10' elements of a sigma70-type promoter. Deletions of the promoters for these genes were analysed with a chloramphenical acetyltransferase (CAT) reporter gene and plasmid constructs that were stably maintained in Escherichia coli or transiently transfected into B. burgdorferi. Reporter expression was measured as susceptibility of transformed E. coli cells to chloramphenical and the CAT activity of E. coli and B. burgdorferi lysates in vitro. Presence of the

'-10' element was essential for full activity in both B. burgdorferi and E. coli. Upstream of the '-35' elements of the ospAB and ***vmp*** promoters were tracts with Ts in 16 of 20 positions for B. burgdorferi and 18 of 20 positions for B. hermsii. Deletion of the T-rich region from the ospAB or ***vmp*** promoter caused a greater reduction of CAT activity in B. burgdorferi than in E. coli. The findings indicate that ospAB and ***vmp*** promoters are extended promoters with two parts: (i) a core region containing typical '-35' and '-10' elements and (ii) a unique T-rich region.

TI The extended promoters for two outer membrane lipoprotein genes of ***Borrelia*** spp. uniquely include a T-rich region.

AB OspA and B proteins of ***Borrelia*** burgdorferi and ***Vmp*** proteins of ***Borrelia*** hermsii are abundant outer membrane lipoproteins, whose expression varies with the environment. The genes for these proteins have the '-35'. . . essential for full activity in both B. burgdorferi and E. coli. Upstream of the '-35' elements of the ospAB and ***vmp*** promoters were tracts with Ts in 16 of 20 positions for B. burgdorferi and 18 of 20 positions for B. hermsii. Deletion of the T-rich region from the ospAB or ***vmp*** promoter caused a greater reduction of CAT activity in B. burgdorferi than in E. coli. The findings indicate that ospAB and ***vmp*** promoters are extended promoters with two parts: (i) a core region containing typical '-35' and '-10' elements and (ii) a. . .

IT . . .

and Molecular Biophysics)

IT Chemicals & Biochemicals

enzymes; outer membrane lipoprotein genes; outer membrane lipoproteins; outer membrane proteins; plasmids; proteins; ***Borrelia*** ospAB gene; ***Borrelia*** ***vmp*** gene

ORGN . . .

Escherichia coli

Taxa Notes

Bacteria, Eubacteria, Microorganisms

ORGN Classifier

Spirochaetaceae 06112

Super Taxa

Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name

Borrelia spp.

Taxa Notes

- L4 ANSWER 43 OF 87 SCISEARCH COPYRIGHT (c) 2009 The Thomson Corporation on $_{\mbox{\scriptsize STN}}$
- AN 1999:121219 SCISEARCH <<LOGINID::20090609>>
- GA The Genuine Article (R) Number: 165HE
- TI Antigenic variation in Lyme disease ***borreliae*** by promiscuous recombination of ***VMP*** -like sequence cassettes (vol 89, pg 275, 1997)
- AU Zhang J R (Reprint)
- CS Univ Texas, Sch Med, Dept Pathol & Lab Med, Houston, TX 77030 USA (Reprint)
- AU Hardham J M; Barbour A G; Norris S J
- CS Univ Texas, Sch Med, Dept Microbiol & Mol Genet, Houston, TX 77030 USA; Univ Calif Irvine, Coll Med, Dept Microbiol & Mol Genet, Irvine, CA 92697 USA
- CYA USA

- SO CELL, (5 FEB 1999) Vol. 96, No. 3, pp. U23-U23. ISSN: 0092-8674.
- PB CELL PRESS, 1100 MASSACHUSETTES AVE,, CAMBRIDGE, MA 02138 USA.
- DT Errata; Journal
- LA English
- REC Reference Count: 1
- ED Entered STN: 1999
 - Last Updated on STN: 1999
- TI Antigenic variation in Lyme disease ***borreliae*** by promiscuous recombination of ***VMP*** -like sequence cassettes (vol 89, pg 275, 1997)
- L4 ANSWER 44 OF 87 CAPLUS COPYRIGHT 2009 ACS on STN
- AN 1999:123920 CAPLUS <<LOGINID::20090609>>
- TI Antigenic variation in lyme disease ***borreliae*** by promiscuous recombination of ***VMP*** -like sequence cassettes
- AU Anon.
- SO Cell (Cambridge, Massachusetts) (1999), 96(3), no pp. Given CODEN: CELLB5; ISSN: 0092-8674
- PB Cell Press
- DT Journal; Errata
- LA English
- AB Unavailable
- TI Antigenic variation in lyme disease ***borreliae*** by promiscuous recombination of ***VMP*** -like sequence cassettes
- L4 ANSWER 45 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN DUPLICATE 26
- AN 1998:393355 BIOSIS <<LOGINID::20090609>>
- DN PREV199800393355
- TI Genetic variation of the ***Borrelia*** burgdorferi gene vlsE involves cassette-specific, segmental gene conversion.
- AU Zhang, Jing-Ren; Norris, Steven J. [Reprint author]
- CS Dep. Pathol. Lab. Med., Univ. Tex. Med. Sch., 6431 Fannin, Houston, TX 77030, USA
- SO Infection and Immunity, (Aug., 1998) Vol. 66, No. 8, pp. 3698-3704. print. CODEN: INFIBR. ISSN: 0019-9567.
- DT Article
- LA English
- ED Entered STN: 10 Sep 1998
 Last Updated on STN: 10 Sep 1998
- The Lyme disease spirochete ***Borrelia*** burgdorferi possesses 15 AB silent ***vls*** cassettes and a ***vls*** expression site (vlsE) encoding a surface-exposed lipoprotein. Segments of the silent ***vls*** cassettes have been shown to recombine with the vlsE cassette region in the mammalian host, resulting in combinatorial antigenic variation. Despite promiscuous recombination within the vlsE cassette region, the 5' and 3' coding sequences of vlsE that flank the cassette region are not subject to sequence variation during these recombination events. The segments of the silent ***vls*** cassettes recombine in the vlsE cassette region through a unidirectional process such that the sequence and organization of the silent ***vls*** loci are not affected. As a result of recombination, the previously expressed segments are replaced by incoming segments and apparently degraded. These results provide evidence for a gene conversion mechanism in VlsE antigenic variation.
- TI Genetic variation of the ***Borrelia*** burgdorferi gene vlsE involves

cassette-specific, segmental gene conversion.

AB The Lyme disease spirochete ***Borrelia*** burgdorferi possesses 15 silent ***vls*** cassettes and a ***vls*** expression site (vlsE) encoding a surface-exposed lipoprotein. Segments of the silent ***vls*** cassettes have been shown to recombine with the vlsE cassette region in the mammalian host, resulting in combinatorial antigenic variation... that flank the cassette region are not subject to sequence variation during these recombination events. The segments of the silent ***vls*** cassettes recombine in the vlsE cassette region through a unidirectional process such that the sequence and organization of the silent ***vls*** loci are not affected. As a result of recombination, the previously expressed segments are replaced by incoming segments and apparently. . .

ORGN Classifier

Spirochaetaceae 06112

Super Taxa

Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name

Borrelia -burgdorferi

Taxa Notes

Bacteria, Eubacteria, Microorganisms

- L4 ANSWER 46 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN DUPLICATE 27
- AN 1998:393354 BIOSIS <<LOGINID::20090609>>
- DN PREV199800393354
- TI Kinetics and in vivo induction of genetic variation of vlsE in ***Borrelia*** burgdorferi.
- AU Zhang, Jing-Ren; Norris, Steven J. [Reprint author]
- CS Dep. Pathol. Lab. Med., Univ. Tex. Med. Sch., 6431 Fannin, Houston, TX 77030, USA
- SO Infection and Immunity, (Aug., 1998) Vol. 66, No. 8, pp. 3689-3697. print. CODEN: INFIBR. ISSN: 0019-9567.
- DT Article
- LA English
- ED Entered STN: 10 Sep 1998

 Last Updated on STN: 10 Sep 1998
- Last Updated on STN: 10 Sep 1998 The Lyme disease agent, ***Borrelia*** burgdorferi, is able to AΒ persistently infect humans and animals for months or years in the presence of an active immune response. It is not known how the organisms survive immune attack in the mammalian host. vlsE, a gene localized near one end of linear plasmid lp28-1 and encoding a surface-exposed lipoprotein in B. burgdorferi B31, was shown recently to undergo extensive genetic and antigenic variation within 28 days of initial infection in C3H/HeN mice. In this study, we examined the kinetics of vlsE sequence variation in C3H/HeN mice at 4, 7, 14, 21, and 28 days and at 7 and 12 months postinfection. Sequence changes were detected by PCR amplification and sequence analysis as early as 4 days postinfection and accumulated progressively in both C3H/HeN and CB-17 severe combined immunodeficient (SCID) mice throughout the course of infection. The sequence changes were consistent with sequential recombination of segments from multiple silent ***vls*** cassette sites into the vlsE expression site. No vlsE sequence changes were detected in organisms cultured in vitro for up to 84 days. These results indicate that vlsE recombination is induced by a factor(s) present in the mammalian host, independent of adaptive immune responses. The possible inducing conditions appear to be present in

various tissue sites because isolates from multiple tissues showed similar

degrees of sequence variation. The rate of accumulation of predicted amino acid changes was higher in the immunologically intact C3H/HeN mice than in SCID mice, a finding consistent with immune selection of VlsE variants.

- TI Kinetics and in vivo induction of genetic variation of vlsE in ***Borrelia*** burgdorferi.
- AB The Lyme disease agent, ***Borrelia*** burgdorferi, is able to persistently infect humans and animals for months or years in the presence of an active immune. . . (SCID) mice throughout the course of infection. The sequence changes were consistent with sequential recombination of segments from multiple silent ***vls*** cassette sites into the vlsE expression site. No vlsE sequence changes were detected in organisms cultured in vitro for up. . .

ORGN . .

Mammals, Nonhuman Vertebrates, Nonhuman Mammals, Rodents, Vertebrates ORGN Classifier

Spirochaetaceae 06112

Super Taxa

Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name

Borrelia -burgdorferi

Taxa Notes

- L4 ANSWER 47 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN DUPLICATE 28
- AN 1998:352289 BIOSIS <<LOGINID::20090609>>
- DN PREV199800352289
- TI Bloodstream-versus tick-associated variants of a relapsing fever bacterium.
- AU Schwan, Tom G. [Reprint author]; Hinnebusch, B. Joseph
- CS Lab. Microbial Structure Function, Rocky Mountain Laboratories, Natl. Inst. Allergy Infectious Diseases, Natl. Inst. Health, Hamilton, MT 59840, USA
- SO Science (Washington D C), (June 19, 1998) Vol. 280, No. 5371, pp. 1938-1940. print. CODEN: SCIEAS. ISSN: 0036-8075.
- DT Article
- LA English
- ED Entered STN: 13 Aug 1998
 Last Updated on STN: 13 Aug 1998
- AB The relapsing fever spirochete, ***Borrelia*** hermsii, alternates infections between a mammal and a tick vector. Whether the spirochete changes phenotypically in the different hosts was examined by allowing the tick vector Ornithodoros hermsi to feed on mice infected with serotype 7 or serotype 8 of B. hermsii. Upon infection of ticks, the spirochetal serotype-specific variable major proteins (***Vmps***) 7 and 8 became undetectable and were replaced by ***Vmp33*** . This switch from a bloodstream- to tick-associated phenotype could be induced in culture by a decrease in temperature. After tick-bite transmission back to mice, the process was reversed and the spirochetes resumed expression of the same ***Vmp*** present in the previous infectious blood meal.
- AB The relapsing fever spirochete, ***Borrelia*** hermsii, alternates infections between a mammal and a tick vector. Whether the spirochete changes phenotypically in the different hosts was. . . infected with serotype 7 or serotype 8 of B. hermsii. Upon infection of ticks, the spirochetal serotype-specific variable major proteins (***Vmps***) 7

and 8 became undetectable and were replaced by ***Vmp33*** . This switch from a bloodstream- to tick-associated phenotype could be induced in culture by a decrease in temperature. After tick-bite transmission back to mice, the process was reversed and the spirochetes resumed expression of the same ***Vmp*** present in the previous infectious blood meal.

ORGN . .

Mammals, Nonhuman Vertebrates, Nonhuman Mammals, Rodents, Vertebrates ORGN Classifier

Spirochaetaceae 06112

Super Taxa

Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name

Borrelia -hermsii: bloodstream variant, pathogen, serovar-7, relapsing fever spirochete, serovar-8, tick-associated variant, tick bite transmission

Taxa Notes

- L4 ANSWER 48 OF 87 CAPLUS COPYRIGHT 2009 ACS on STN DUPLICATE 29
- AN 1998:787140 CAPLUS <<LOGINID::20090609>>
- DN 130:152425
- TI Variable major lipoprotein is a principal TNF-inducing factor of louse-borne relapsing fever
- AU Vidal, Vincent; Scragg, Ian G.; Cutler, Sally J.; Rockett, Kirk A.; Rekade, Daniel; Warrell, David A.; Wright, David J. M.; Kwiatkowski,
- CS University Department of Pediatrics, Oxford University, UK
- SO Nature Medicine (New York) (1998), 4(12), 1416-1420 CODEN: NAMEFI; ISSN: 1078-8956
- PB Nature America
- DT Journal
- LA English
- AΒ Massive release of tumor necrosis factor is responsible for the potentially fatal Jarisch-Herxheimer reaction that follows antibiotic treatment of relapsing fever due to ***Borrelia*** recurrentis. The authors have undertaken the quant. purifn. of the components of B. recurrentis that stimulate human monocytes to produce tumor necrosis factor. The authors show that the predominant factor inducing tumor necrosis factor is a variable lipoprotein homologous to the variable major protein of B. hermsii. The authors found antibodies to different forms of variable major protein in two patients with louse-borne relapsing fever. The three purified variable major proteins studied here differ in their ability to induce tumor necrosis factor prodn., which may partly explain ***borrelial*** infection. These the variable clin. severity of results may be of considerable relevance for the pathogenesis of Lyme disease and other forms of human ***borreliosis***
- RE.CNT 27 THERE ARE 27 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- AB . . . tumor necrosis factor is responsible for the potentially fatal Jarisch-Herxheimer reaction that follows antibiotic treatment of relapsing fever due to ***Borrelia*** recurrentis. The authors have undertaken the quant. purifn. of the components of B. recurrentis that stimulate human monocytes to produce. . . here differ in their ability to induce tumor necrosis factor prodn., which may partly explain the variable clin. severity of ***borrelial*** infection. These results may be of considerable relevance for the pathogenesis of Lyme disease and other

borreliosis . forms of human ***vmpA1*** lipoprotein tumor necrosis factor louse borne relapsing ST fever; sequence variable major protein ***vmpA1*** ***Borrelia*** ΤТ Disease, animal (Jarisch-Herxheimer reaction; variable major lipoprotein A1 of ***Borrelia*** recurrentis sequence and as principal tumor necrosis factor-inducing factor of louse-borne relapsing fever in humans in relation to) ΙT Lipoproteins RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study) (***VmpA1*** (variable major protein A1); variable major lipoprotein A1 of ***Borrelia*** recurrentis sequence and as principal tumor necrosis factor-inducing factor of louse-borne relapsing fever in humans) ΙT Fever and Hyperthermia (louse-borne relapsing; variable major lipoprotein A1 of ***Borrelia*** recurrentis sequence and as principal tumor necrosis factor-inducing factor of louse-borne relapsing fever in humans) ΙT DNA sequences (of variable major protein A1 gene ***vmpA1*** of ***Borrelia*** recurrentis) Protein sequences ΙT (of variable major protein Al of ***Borrelia*** recurrentis) ***Borrelia*** recurrentis ΤT (variable major lipoprotein A1 of ***Borrelia*** recurrentis sequence and as principal tumor necrosis factor-inducing factor of louse-borne relapsing fever in humans) Tumor necrosis factors ΤТ RL: BAC (Biological activity or effector, except adverse); BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process) (variable major lipoprotein Al of ***Borrelia*** recurrentis sequence and as principal tumor necrosis factor-inducing factor of louse-borne relapsing fever in humans) ΙT Gene, microbial RL: PRP (Properties) (***vmpA1*** ; variable major lipoprotein A1 of ***Borrelia*** recurrentis sequence and as principal tumor necrosis factor-inducing factor of louse-borne relapsing fever in humans) 206631-77-8, GenBank AJ224157 ΤТ RL: PRP (Properties) (nucleotide sequence; variable major lipoprotein A1 of ***Borrelia*** recurrentis sequence and as principal tumor necrosis factor-inducing factor of louse-borne relapsing fever in humans) ANSWER 49 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on L4DUPLICATE 30 1998:120628 BIOSIS <<LOGINID::20090609>> ΑN DNPREV199800120628 ΤI Population structure of the relapsing fever spirochete ***Borrelia*** hermsii as indicated by polymorphism of two multigene families that encode immunogenic outer surface lipoproteins. ΑU Hinnebusch, B. Joseph [Reprint author]; Barbour, Alan G.; Restrepo, Blanca

Lab. Microbial Structure Function, Rocky Mountain Lab., Natl. Inst.

I.; Schwan, Tom G.

CS

Allergy Infectious Diseases, Natl. Inst. Health, 903 S. 4th St., Hamilton, MT 59840, USA

- SO Infection and Immunity, (Feb., 1998) Vol. 66, No. 2, pp. 432-440. print. CODEN: INFIBR. ISSN: 0019-9567.
- DT Article
- LA English
- ED Entered STN: 5 Mar 1998

 Last Updated on STN: 5 Mar 1998
- AΒ The tick-borne relapsing fever spirochete ***Borrelia*** evades the mammalian immune system by periodically switching expression among members of two multigene families that encode immunogenic, antigenically distinct outer surface proteins. The type strain, B. hermsii HS1, has at least 40 complete genes and pseudogenes that participate in this multiphasic antigenic variation. Originally termed ***vmp*** (for variable major protein) genes, they have been reclassified as vsp (for variable small protein) and vlp (for variable large protein) genes, based on size and amino acid sequence similarities. To date, antigenic variation in B. hermsii has been studied only in the type strain, HS1. Nucleotide sequence comparisons of 23 B. hermsii HS1 genes revealed five distinct groups, the vsp gene family and four subfamilies of vlp genes. We used PCR with family- and subfamily-specific primers, followed by restriction fragment length polymorphism analysis, to compare the vsp and vlp repertoires of HS1 and seven other B. hermsii isolates from Washington, Idaho, and California. This analysis, together with pulsed-field gel electrophoresis genome profiles, revealed that the eight isolates formed three distinct groups, which likely represent clonal lineages. Members of the three groups coexisted in the same geographic area, but they could also be isolated across large geographical distances. This population structure may result from immune selection by the host, as has been proposed for other pathogens with polymorphic antigens.
- TI Population structure of the relapsing fever spirochete ***Borrelia***
 hermsii as indicated by polymorphism of two multigene families that encode
 immunogenic outer surface lipoproteins.
- AB The tick-borne relapsing fever spirochete ***Borrelia*** hermsii evades the mammalian immune system by periodically switching expression among members of two multigene families that encode immunogenic, antigenically. . . B. hermsii HS1, has at least 40 complete genes and pseudogenes that participate in this multiphasic antigenic variation. Originally termed ***vmp*** (for variable major protein) genes, they have been reclassified as vsp (for variable small protein) and vlp (for variable large. . .

ORGN Classifier

Spirochaetaceae 06112

Super Taxa

Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name

Borrelia -hermsii: tick-borne relapsing fever spirochete Taxa Notes

- L4 ANSWER 50 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN
- AN 1998:416557 BIOSIS <<LOGINID::20090609>>
- DN PREV199800416557
- TI Human tissue culture cells induce the ***Vmp*** -like outer membrane protein, VlsE, in ***Borrelia*** burgdorferi.
- AU Frye, J. G.; Hudson, C. R.; Gherardini, F. C.

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CS
    Univ. Ga., Athens, GA, USA
    Abstracts of the General Meeting of the American Society for Microbiology,
SO
     (1998) Vol. 98, pp. 222. print.
    Meeting Info.: 98th General Meeting of the American Society for
    Microbiology. Atlanta, Georgia, USA. May 17-21, 1998. American Society for
    Microbiology.
    ISSN: 1060-2011.
DT
    Conference; (Meeting)
    Conference; Abstract; (Meeting Abstract)
    Conference; (Meeting Poster)
LA
    English
ED
    Entered STN: 2 Oct 1998
    Last Updated on STN: 2 Oct 1998
ΤI
    Human tissue culture cells induce the ***Vmp*** -like outer membrane
    protein, VlsE, in ***Borrelia*** burgdorferi.
    . . .
ΙT
       Bacteriology; Genetics
ΙT
    Diseases
       Lyme disease: bacterial disease
       Lyme Disease (MeSH)
    Chemicals & Biochemicals
ΙT
       mRNA [messenger RNA]; DNA; RNA; VlsE: ***Vmp*** -like outer membrane
       protein
ORGN .
Notes
       Animals, Chordates, Humans, Mammals, Primates, Vertebrates
ORGN Classifier
       Spirochaetaceae 06112
     Super Taxa
        Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms
     Organism Name
           ***Borrelia*** -burgdorferi: pathogen
       Bacteria, Eubacteria, Microorganisms
    ANSWER 51 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on
L4
                                                       DUPLICATE 31
    1998:213424 BIOSIS <<LOGINID::20090609>>
ΑN
    PREV199800213424
DN
    Genetic and immunological analyses of VIs ( ***VMP*** -like sequences)
TΙ
          ***Borrelia*** burgdorferi.
    Kawabata, Hiroki; Myouga, Fumiyoshi; Inagaki, Yoshishige; Murai, Noriyuki;
ΑU
    Watanabe, Haruo [Reprint author]
    Dep. Bacteriol., Natl. Inst. Infect. Dis., 1-23-1 Toyama, Shinjyuku-ku,
CS
    Tokyo 162, Japan
SO
    Microbial Pathogenesis, (March, 1998) Vol. 24, No. 3, pp. 155-166. print.
    CODEN: MIPAEV. ISSN: 0882-4010.
DT
    Article
    English
LA
    Entered STN: 11 May 1998
ED
    Last Updated on STN: 11 May 1998
    DNA fragments containing the ***VMP*** -like sequence ( ***Vls***
AΒ
    were cloned from ***Borrelia*** burgdorferi strain 297. Analyses by
    PCR, PFGE, and Southern hybridization revealed that the ***Vls***
     sequences existed in multi-copies on the 20-kb ***borrelial***
    plasmid, but not on chromosomes or other plasmids. One
                                                             ***Vls*** unit
    of the strain 297 was about 669 bases, and predicted peptides length was
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223 amino acids. Homologues of the ***Vls*** fragment were detected
     in three B. burgdorferi strains, a B. garinii strain 20047, and a B.
     afzelii strain P/Gau. A recombinant VlsII protein prepared in Escherichia
     coli strain JM109 reacted with antibodies that existed in three of five
     patients, by immunoblotting. These results suggested that the ***Vls***
     of B. burgdorferi is expressed in Lyme disease patients.
    Genetic and immunological analyses of VIs ( ***VMP*** -like sequences)
    of ***Borrelia*** burgdorferi.
    DNA fragments containing the ***VMP*** -like sequence ( ***Vls*** )
    were cloned from ***Borrelia*** burgdorferi strain 297. Analyses by
    PCR, PFGE, and Southern hybridization revealed that the
                                                            ***Vls***
     sequences existed in multi-copies on the 20-kb ***borrelial***
    plasmid, but not on chromosomes or other plasmids. One ***Vls***
    of the strain 297 was about 669 bases, and predicted peptides length was
     223 amino acids. Homologues of the ***Vls*** fragment were detected
     in three B. burgdorferi strains, a B. garinii strain 20047, and a B.
     afzelii strain P/Gau. A. . . coli strain JM109 reacted with antibodies
    that existed in three of five patients, by immunoblotting. These results
     suggested that the ***Vls*** of B. burgdorferi is expressed in Lyme
    disease patients.
Diseases
       Lyme disease: bacterial disease
       Lyme Disease (MeSH)
    Chemicals & Biochemicals
       ospC gene; outer surface protein C; variable major protein; ***vls***
ORGN Classifier
       Spirochaetaceae
                        06112
     Super Taxa
       Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms
    Organism Name
           ***Borrelia*** -burgdorferi: pathogen
           ***Borrelia*** -hermsii: pathogen
     Taxa Notes
       Bacteria, Eubacteria, Microorganisms
    ANSWER 52 OF 87 CAPLUS COPYRIGHT 2009 ACS on STN
    1998:152421 CAPLUS <<LOGINID::20090609>>
    128:201751
OREF 128:39807a,39810a
    Antigenic variation in Lyme disease spirochetes by segmental recombination
         ***vmp*** -like sequence cassettes
     Zhang, Jing-Ren
    Health Science Center, Univ. of Texas, Houston, TX, USA
    (1997) 126 pp. Avail.: UMI, Order No. DA9813074
    From: Diss. Abstr. Int., B 1998, 58(10), 5258
    Dissertation
    English
    Unavailable
    Antigenic variation in Lyme disease spirochetes by segmental recombination
    of ***vmp*** -like sequence cassettes
                                                               ***vmp***
    antiquenic variation Lyme disease spirochete recombination;
    cassette sequence segmental recombination ***Borrelia***
    Antigenic variation
        ***Borrelia*** burgdorferi
     Recombination, genetic
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AB

ΙT

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(antigenic variation in Lyme disease spirochetes by segmental recombination of ***vmp*** -like sequence cassettes) ΙT Antigens RL: BOC (Biological occurrence); BSU (Biological study, unclassified); BIOL (Biological study); OCCU (Occurrence) (antigenic variation in Lyme disease spirochetes by segmental recombination of ***vmp*** -like sequence cassettes) ΤТ Gene, microbial RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process) (***vmp*** ; antigenic variation in Lyme disease spirochetes by segmental recombination of ***vmp*** -like sequence cassettes) L4ANSWER 53 OF 87 CAPLUS COPYRIGHT 2009 ACS on STN ΑN 1997:579836 CAPLUS <<LOGINID::20090609>> 127:189742 DN OREF 127:36809a,36812a ΤI ***Vmp*** -like sequences of pathogenic ***Borrelia*** ΙN Norris, Steven J.; Zhang, Jing-ren; Hardham, John M.; Howell, Jerrilyn K.; Barbour, Alan G.; Weinstock, George M. Board of Regents, the University of Texas System, USA; Norris, Steven J.; PAZhang, Jing-Ren; Hardham, John M.; Howell, Jerrilyn K.; Barbour, Alan G.; Weinstock, George M. PCT Int. Appl., 130 pp. SO CODEN: PIXXD2 DT Patent English FAN.CNT 1 KIND DATE APPLICATION NO. DATE PATENT NO. _____ ____ _____ _____ A1 19970828 WO 1997-US2952 19970220 WO 9731123 W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, HU, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU RW: KE, LS, MW, SD, SZ, UG, AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG AU 9721915 A 19970910 AU 1997-21915 19970220 A1 19990203 EP 1997-914794 EP 894143 19970220 B1 20050810 EP 894143 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI AT 1997-914794 20050815 AT 301716 Τ 19970220 19970220 EP 1589109 Α2 20051026 EP 2005-10338 EP 1589109 А3 20051116 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI US 6437116 В1 20020820 US 1999-125619 19990127 US 20030092903 A1 US 2002-143024 20030515 20020731 В2 US 6740744 20040525 US 20030060618 A1 20030327 US 2002-222162 20020816
US 6878816 B2 20050412
US 20040044192 A1 20040304 US 2002-222566 20020816
US 6719983 B2 20040413
US 20040214225 A1 20041028 US 2004-852555 20040524
US 7135176 B2 20061114

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US 20070117970
                       A1
                             20070524 US 2006-501166 20060807
PRAI US 1996-12028P
                        Р
                               19960221
    EP 1997-914794
                        A3
                               19970220
    WO 1997-US2952
                        W
                               19970220
    US 1999-125619
                        А3
                               19990127
    US 2002-143024
                        А3
                               20020731
     US 2002-222162
                        А3
                               20020816
                               20040524
    US 2004-852555
                        А3
    The present invention relates to DNA sequences encoding ***Vmp*** -like
AB
    polypeptides of pathogenic ***Borrelia*** , the use of the DNA
     sequences in recombinant vectors to express polypeptides, the encoded
     amino acid sequences, application of the DNA and amino acid sequences to
     the prodn. of polypeptides as antigens for immunoprophylaxis,
     immunotherapy, and immunodiagnosis. Also disclosed are the use of the
    nucleic acid sequences as probes or primers for the detection of organisms
     causing Lyme disease, relapsing fever, or related disorders, and kits
     designed to facilitate methods of using the described polypeptides, DNA
     segments and antibodies.
             THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT 1
             ALL CITATIONS AVAILABLE IN THE RE FORMAT
ΤI
      ***Vmp*** -like sequences of pathogenic ***Borrelia***
                                                            ***Vmp*** -like
AΒ
    The present invention relates to DNA sequences encoding
    polypeptides of pathogenic ***Borrelia*** , the use of the DNA
     sequences in recombinant vectors to express polypeptides, the encoded
    amino acid sequences, application of the. .
    variable major protein gene
                                 ***Borrelia***
ST
ΙT
      ***Borrelia*** burgdorferi
        ( ***Vmp*** -like sequences of pathogenic ***Borrelia*** )
ΙT
    Proteins, specific or class
     RL: BOC (Biological occurrence); BSU (Biological study, unclassified);
     BIOL (Biological study); OCCU (Occurrence)
        (gene vlsE; ***Vmp*** -like sequences of pathogenic ***Borrelia***
    189614-97-9, DNA ( ***Borrelia*** burgdorferi strain B31 clone 5A3 gene
ΙT
    vlsE plus flanks)
     RL: BOC (Biological occurrence); BSU (Biological study, unclassified);
     BIOL (Biological study); OCCU (Occurrence)
       ( ***Vmp*** -like sequences of pathogenic ***Borrelia*** )
    189833-73-6, Protein ( ***Borrelia*** burgdorferi strain B31 clone 5A3
ΤТ
     gene vlsE)
     RL: BOC (Biological occurrence); BSU (Biological study, unclassified); THU
     (Therapeutic use); BIOL (Biological study); OCCU (Occurrence); USES (Uses)
        ( ***Vmp*** -like sequences of pathogenic ***Borrelia*** )
L4
    ANSWER 54 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on
                                                      DUPLICATE 32
    1997:391165 BIOSIS <<LOGINID::20090609>>
ΑN
    PREV199799690368
    Immunologic and genetic analyses of ***VmpA*** of a neurotropic strain
TΤ
        ***Borrelia*** turicatae.
ΑU
    Cadavid, Diego; Pennington, Pamela M.; Kerentseva, Tatiana A.; Bergstrom,
     Sven; Barbour, Alan G. [Reprint author]
CS
    Dep. Microbiol. Molecular Genetics, Univ. California Irvine, Irvine, CA
    92697-4025, USA
SO
    Infection and Immunity, (1997) Vol. 65, No. 8, pp. 3352-3360.
    CODEN: INFIBR. ISSN: 0019-9567.
DT
    Article
```

LA English

ED Entered STN: 10 Sep 1997
Last Updated on STN: 10 Sep 1997

AΒ In mice infected with serotype A but not serotype B of the relapsing fever ***Borrelia*** turicatae, early invasion of the brain spirochete occurs. Serotypes A and B are further distinguished by the abundant surface protein they produce: ***VmpA*** and ***UmpB*** respectively. Western blotting with monoclonal antibodies, one-dimensional peptide mapping, and partial amino acid sequencing demonstrated regions of the ***VmpA*** protein that differed from ***VmpB*** . Oligonucleotide primers based on the partial amino acid sequences of unique regions were used to amplify a portion of the ***VmpA*** gene (***vmpA***) by PCR, and the product was used as a probe in Southern blot and Northern blot analyses. These experiments showed that (i) expression of the ***vmpA*** sequence was determined at the level of transcription and (ii) the ***vmpA*** sequence was in two locations in serotype A and one location in serotype B. The $\mbox{\tt ***vmpA***}$ gene at the expression-linked locus of serotype A was cloned and sequenced. An open reading frame would encode a polypeptide of 214 amino acids. The polypeptide expressed by Escherichia coli was bound by VmA-specific but not ***VmpB*** -specific antibody. Primer extension analysis identified a consensus sigma-70-type promoter for ***vmpA*** at the expression locus. Phylogenetic analysis revealed that ***VmpA*** is homologous to small ***Vmp*** (Vsp) proteins of B. hermsii and to OspC proteins of B. burgdorferi. These findings indicate that a function of the Vsp-OspC family of proteins of ***Borrelia*** spp. may be differential localization in organs, including the brain, during infection.

TI Immunologic and genetic analyses of ***VmpA*** of a neurotropic strain of ***Borrelia*** turicatae.

In mice infected with serotype A but not serotype B of the relapsing fever AΒ spirochete ***Borrelia*** turicatae, early invasion of the brain occurs. Serotypes A and B are further distinguished by the abundant surface protein they produce: ***VmpA*** and ***VmpB*** respectively. Western blotting with monoclonal antibodies, one-dimensional peptide mapping, and partial amino acid sequencing demonstrated regions of the ***VmpA*** protein that differed from ***VmpB*** . Oligonucleotide primers based on the partial amino acid sequences of unique regions were used to amplify a portion of the ***VmpA*** gene (***vmpA***) by PCR, and the product was used as a probe in Southern blot and Northern blot analyses. These experiments showed that (i) expression of the ***vmpA*** sequence was determined at the level of transcription and (ii) the ***vmpA*** sequence was in two locations in serotype A and one location in serotype B. The ***vmpA*** gene at the expression-linked locus of serotype A was cloned and sequenced. An open reading frame would encode a polypeptide of 214 amino acids. The polypeptide expressed by Escherichia coli was bound by VmA-specific but not ***VmpB*** -specific antibody. Primer extension analysis identified a consensus sigma-70-type promoter for ***vmpA*** at the expression locus. Phylogenetic analysis revealed that ***VmpA*** is homologous to small ***Vmp*** (Vsp) proteins of B. hermsii and to OspC proteins of B. burgdorferi. These findings indicate that a function of the Vsp-OspC family of proteins of ***Borrelia*** spp. may be differential localization in organs, including the brain, during infection.

IT . . .

GENETICS; NCBI DATABANK; NERVOUS SYSTEM DISEASE; POLYMERASE CHAIN REACTION; SEROVAR-A; SEROVAR-B; STRUCTURE; U85413; ***VMPA*** GENE ORGN Classifier Spirochaetaceae 06112 Super Taxa Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name ***Borrelia*** turicatae Taxa Notes Bacteria, Eubacteria, Microorganisms L4ANSWER 55 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on DUPLICATE 33 ΑN 1997:225972 BIOSIS <<LOGINID::20090609>> DMPREV199799517688 Antigenic variation in Lyme disease ***Borreliae*** by promiscuous ТΤ recombination of ***VMP*** -like sequence cassettes. ΑU Zhang, Jing-Ren [Reprint author]; Hardham, John M.; Barbour, Alan G.; Norris, Steven J. CS Dep. Pathol., Univ. Texas Med. Sch. Houston, Houston, TX 77030, USA SO Cell, (1997) Vol. 89, No. 2, pp. 275-285. CODEN: CELLB5. ISSN: 0092-8674. Article DT Enalish LA ED Entered STN: 22 May 1997 Last Updated on STN: 22 May 1997 AB We have identified and characterized an elaborate genetic system in the Lyme disease spirochete ***Borrelia*** burgdorferi that promotes extensive antigenic variation of a surface-exposed lipoprotein, VlsE. A 28 kb linear plasmid of B. burgdorferi B31 (lp28-1) was found to contain a ***vmp*** -like sequence (***vls***) locus that closely resembles the variable major protein (***vmp***) system for antigenic variation of relapsing fever organisms. Portions of several of the 15 nonexpressed (silent) ***vls*** cassette sequences located upstream of vlsE recombined into the central vlsE cassette region during infection of C3H/HeN mice, resulting in antigenic variation of the expressed lipoprotein. This combinatorial variation could potentially produce millions of antigenic variants in the mammalian host. Antigenic variation in Lyme disease ***Borreliae*** by promiscuous TΙ recombination of ***VMP*** -like sequence cassettes. We have identified and characterized an elaborate genetic system in the AB Lyme disease spirochete ***Borrelia*** burgdorferi that promotes extensive antiquenic variation of a surface-exposed lipoprotein, VlsE. A 28 kb linear plasmid of B. burgdorferi B31 (lp28-1) was found to contain a ***vmp*** -like sequence (***vls***) locus that closely resembles the variable major protein (***vmp***) system for antigenic variation of relapsing fever organisms. Portions of several of the 15 nonexpressed (silent) ***vls*** cassette sequences located upstream of vlsE recombined into the central vlsE cassette region during infection of C3H/HeN mice, resulting in. . TΤ Miscellaneous Descriptors ANTIGENIC VARIATION; BACTERIAL DISEASE; COMBINATORIAL VARIATION; C3H/HEN; INFECTION; LYME DISEASE; MOLECULAR GENETICS; PATHOGEN; PROMISCUOUS RECOMBINATION; SURFACE-EXPOSED LIPOPROTEIN; VLSE;

VMP -LIKE SEQUENCE CASSETTES

ORGN . . . Mammals, Nonhuman Vertebrates, Nonhuman Mammals, Rodents, Vertebrates ORGN Classifier Spirochaetaceae 06112 Super Taxa Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name ***Borrelia*** burgdorferi Taxa Notes Bacteria, Eubacteria, Microorganisms L4ANSWER 56 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on ΑN 1997:282165 BIOSIS <<LOGINID::20090609>> DN PREV199799581368 Antigenic variation in Lyme disease spirochetes by promiscuous TΤ recombination of ***vmp*** -like sequence cassettes. ΑU Zhang, Jing-Ren [Reprint author]; Hardham, John M.; Barbour, Alan G.; Norris, Steven J. CS Dep. Pathol. Lab. Med., Univ. Texas Med. Sch., Houston, TX, USA Abstracts of the General Meeting of the American Society for Microbiology, SO (1997) Vol. 97, No. 0, pp. 103. Meeting Info.: 97th General Meeting of the American Society for Microbiology. Miami Beach, Florida, USA. May 4-8, 1997. ISSN: 1060-2011. DT Conference; (Meeting) Conference; Abstract; (Meeting Abstract) Conference; (Meeting Poster) LA English Entered STN: 3 Jul 1997 ΕD Last Updated on STN: 3 Jul 1997 Antigenic variation in Lyme disease spirochetes by promiscuous recombination of ***vmp*** -like sequence cassettes. Mammals, Nonhuman Vertebrates, Nonhuman Mammals, Rodents, Vertebrates ORGN Classifier Spirochaetaceae 06112 Super Taxa Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name ***Borrelia*** burgdorferi Taxa Notes Bacteria, Eubacteria, Microorganisms L4ANSWER 57 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN DUPLICATE 34 2002:51422 BIOSIS <<LOGINID::20090609>> ΑN PREV200200051422 Cloning and expression of soluble truncated variants of ***Borrelia*** TΤ OSPA, OSPB and ***VMP7*** ΑU Dunn, J. J. [Inventor]; Barbour, A. G. [Inventor] Bellport, N.Y., USA CS ASSIGNEE: ASSOCIATED UNIVERSITIES, INC. PΤ US 5571718 19961105 Official Gazette of the United States Patent and Trademark Office Patents,

(Nov. 5, 1996) Vol. 1192, No. 1, pp. 420. print.

CODEN: OGUPE7. ISSN: 0098-1133.

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DT
    Patent
LA
    English
ED
    Entered STN: 2 Jan 2002
    Last Updated on STN: 25 Feb 2002
    Cloning and expression of soluble truncated variants of ***Borrelia***
TΤ
    OSPA, OSPB and ***VMP7***
T. 4
    ANSWER 58 OF 87 CAPLUS COPYRIGHT 2009 ACS on STN
    1996:583427 CAPLUS <<LOGINID::20090609>>
ΑN
DN
    125:266978
OREF 125:49633a,49636a
    Homology of variable major protein genes between ***Borrelia***
    hermsii and ***Borrelia*** miyamotoi. [Erratum to document cited in
    CA125:159805]
ΑU
    Hamase, Akiko; Takahashi, Yukie; Nohgi, Keiko; Fukunaga, Masahito
    Laboratory of Molecular Microbiology, Faculty of Pharmacy and
CS
    Pharmaceutical Sciences, Fukuyama University, Hiroshima, 729-02, Japan
SO
    FEMS Microbiology Letters (1996), 143(2-3), 299
    CODEN: FMLED7; ISSN: 0378-1097
PΒ
    Elsevier
    Journal
DT
    English
LA
    The errors were not reflected in the abstr. or the index entries.
AB
    Homology of variable major protein genes between ***Borrelia***
ΤI
    hermsii and ***Borrelia*** miyamotoi. [Erratum to document cited in
    CA125:159805]
                               ***vmp*** like gene DNA; ***Borrelia***
ST
    erratum ***Borrelia***
      ***vmp*** like gene DNA erratum; ***vmp*** like gene DNA sequence
     erratum; variable major protein sequence ***Borrelia*** erratum
ΙT
    Proteins, specific or class
     RL: BOC (Biological occurrence); BSU (Biological study, unclassified); PRP
     (Properties); BIOL (Biological study); OCCU (Occurrence)
        ( ***VMP*** (variable major proteins); characterization of
         ***Borrelia*** miyamotoi ***vmp*** -like genes from strains HT31
       and FR64b and ***Borrelia*** hermsii and comparison of amino acid
       sequence to published ***vmp*** proteins of ***Borrelia***
       hermsii (Erratum))
      ***Borrelia*** hermsii
ΙT
        ***Borrelia*** miyamotoi
     Deoxyribonucleic acid sequences
     Protein sequences
        (characterization of ***Borrelia*** miyamotoi ***vmp*** -like
       genes from strains HT31 and FR64b and ***Borrelia*** hermsii and
       comparison of amino acid sequence to published ***vmp*** proteins
       of ***Borrelia*** hermsii (Erratum))
ΤТ
    Gene, microbial
    RL: BOC (Biological occurrence); BSU (Biological study, unclassified); PRP
     (Properties); BIOL (Biological study); OCCU (Occurrence)
        ( ***vmp*** -like; characterization of ***Borrelia***
                                                                 miyamotoi
         ***vmp*** -like genes from strains HT31 and FR64b and
***Borrelia***
       hermsii and comparison of amino acid sequence to published ***vmp***
       proteins of ***Borrelia*** hermsii (Erratum))
ΤТ
     180291-11-6
     RL: BOC (Biological occurrence); BSU (Biological study, unclassified); PRP
     (Properties); BIOL (Biological study); OCCU (Occurrence)
        (amino acid sequence; characterization of ***Borrelia*** miyamotoi
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vmp -like genes from strains HT31 and FR64b and ***Borrelia*** hermsii and comparison of amino acid sequence to published ***qmv*** proteins of ***Borrelia*** hermsii (Erratum)) 180291-12-7 180308-88-7 ΙT RL: BOC (Biological occurrence); BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study); OCCU (Occurrence) (characterization of ***Borrelia*** miyamotoi ***vmp*** -like genes from strains HT31 and FR64b and ***Borrelia*** hermsii and comparison of amino acid sequence to published ***vmp*** of ***Borrelia*** hermsii (Erratum)) 180308-87-6 ΙT RL: BOC (Biological occurrence); BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study); OCCU (Occurrence) (nucleotide sequence; characterization of ***Borrelia*** ***vmp*** -like genes from strains HT31 and FR64b and ***Borrelia*** hermsii and comparison of amino acid sequence to published ***vmp*** proteins of ***Borrelia*** hermsii (Erratum)) ANSWER 59 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on L4DUPLICATE 35 1996:379860 BIOSIS <<LOGINID::20090609>> ΑN PREV199699102216 DΝ ТΤ Homology of variable major protein genes between ***Borrelia*** hermsii and ***Borrelia*** miyamotoi. ΑU Hamase, Akiko; Takahashi, Yukie; Nohgi, Keiko; Fukunaga, Masahito [Reprint author] CS Lab. Mol. Microbiol., Fac. Pharmacy Pharmaceutical Sci., Fukuyama Univ., Sanzo 985, Japan FEMS Microbiology Letters, (1996) Vol. 140, No. 2-3, pp. 131-137. SO CODEN: FMLED7. ISSN: 0378-1097. DT Article LA English ED Entered STN: 26 Aug 1996 Last Updated on STN: 26 Aug 1996 AΒ Antigenic variation has been studied in detail for the etiological agent of relapsing fever, ***Borrelia*** hermsii. The variable major proteins (***vmps***) are found at its cell surface, enabling it to avoid the host's immune response. We have cloned and sequenced the ***vmp*** -qene (***vmp***)-like sequences from the ***Borrelia*** miyamotoi strains HT31 and FR64b and the deduced amino acid sequences were compared with the published ***vmp*** proteins ***vmp3*** , ***vmp24*** , and ***vmp33*** of B. hermsii. The sequences were aligned and revealed pairwise sequence identities ranging from 45 to 51%, and differences were scattered throughout the sequences. Southern hybridization using the cloned ***vmp*** -like sequence of strain HT31 as a probe suggested that the ump homologues reside on the linear plasmids of B. miyamotoi. The probe hybridized weakly with B. hermsii linear plasmids and restriction digests. These results suggest that B. miyamotoi has sequences resembling the ump genes in B. hermsii. ΤI Homology of variable major protein genes between ***Borrelia***

Antigenic variation has been studied in detail for the etiological agent of relapsing fever, ***Borrelia*** hermsii. The variable major proteins (***vmps***) are found at its cell surface, enabling it to avoid the host's immune response. We have cloned and sequenced the

hermsii and ***Borrelia*** miyamotoi.

AΒ

```
***vmp*** -gene ( ***vmp*** )-like sequences from the ***Borrelia***
    miyamotoi strains HT31 and FR64b and the deduced amino acid sequences were
     compared with the published ***vmp*** proteins ***vmp3*** ,
       ***vmp24*** , and ***vmp33*** of B. hermsii. The sequences were
     aligned and revealed pairwise sequence identities ranging from 45 to 51%,
     and differences were scattered throughout the sequences. Southern
     hybridization using the cloned ***vmp*** -like sequence of strain HT31
     as a probe suggested that the ump homologues reside on the linear plasmids
     of B. miyamotoi.. .
ORGN Classifier
       Spirochaetaceae 06112
     Super Taxa
       Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms
     Organism Name
           ***Borrelia***
                           hermsii
            ***Borrelia*** miyamotoi
     Taxa Notes
       Bacteria, Eubacteria, Microorganisms
    ANSWER 60 OF 87 CABA COPYRIGHT 2009 CABI on STN
    97:98689 CABA <<LOGINID::20090609>>
    19970502811
    The ***Borrelia*** turicatae murine model of Lyme disease
    Barbour, A. G.
    Department of Medicine (Infectious Disease) and Microbiology, University
     of Texas Health Science Center, San Antonio, TX 78284, USA.
    Journal of Spirochetal and Tick-borne Diseases, (1996) Vol. 3, No. 1, pp.
    62-66. 19 ref.
    Journal
    English
    Entered STN: 16 Sep 1997
    Last Updated on STN: 16 Sep 1997
      ***Borrelia*** turicatae is an agent of relapsing fever. During
    relapsing fever, spirochaetes avoid the immune response of the host by a
    multiphasic antigenic variation. In B. hermsii, another agent of relapsing
     fever, the mechanism for the switch in antigens is a gene rearrangement,
     namely an interplasmidic gene conversion or an intraplasmidic deletion
     between direct repeats. In severe combined immunodeficiency (scid) mice,
     B. turicatae causes the constellation of arthritis, myocarditis, uveitis
     and a cranial nerve disorder. In this way, the infection in these mice is
     similar to Lyme disease. Moreover, B. turicatae invades and persists in
    the central nervous system of laboratory mice. The severity of illness,
    particularly the arthritis, and the entry into the brain appear to be
    determined by the small ***Vmp*** proteins of this species. These
    proteins are homologous to the polymorphic OspC proteins of B.
    burgdorferi, the agent of Lyme disease.
    The ***Borrelia*** turicatae murine model of Lyme disease.
       ***Borrelia*** turicatae is an agent of relapsing fever. During
    relapsing fever, spirochaetes avoid the immune response of the host by a.
     . . The severity of illness, particularly the arthritis, and the entry
     into the brain appear to be determined by the small ***Vmp*** proteins
     of this species. These proteins are homologous to the polymorphic OspC
     proteins of B. burgdorferi, the agent of Lyme. . .
      ***Borrelia***; Spirochaetaceae; Spirochaetales; Gracilicutes;
    bacteria; prokaryotes; Muridae; rodents; mammals; vertebrates; Chordata;
     animals; small mammals
     ***Borrelia*** hermsii; ***Borrelia*** burgdorferi;
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- L4 ANSWER 61 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN DUPLICATE 36
- AN 1995:320318 BIOSIS <<LOGINID::20090609>>
- DN PREV199598334618
- TI Evolution of the ***Borrelia*** burgdorferi outer surface protein OspC.
- AU Theisen, Michael [Reprint author]; Borre, Martin; Mathiesen, Marianne J.; Mikkelsen, Bo; Lebech, Anne-Mette; Hansen, Klaus
- CS Dep. Infection-Immunol., Statens Seruminstitut, Artillerivej 5, Copenhagen DK-2300, Denmark
- SO Journal of Bacteriology, (1995) Vol. 177, No. 11, pp. 3036-3044. CODEN: JOBAAY. ISSN: 0021-9193.
- DT Article
- LA English
- ED Entered STN: 30 Jul 1995 Last Updated on STN: 30 Jul 1995
- The genes coding for outer surface protein OspC from 22 ***Borrelia*** burgdorferi strains isolated from patients with Lyme ***borreliosis*** were cloned and sequenced. For reference purposes, the 16S rRNA genes from 17 of these strains were sequenced after being cloned. The deduced OspC amino acid sequences were aligned with 12 published OspC sequences and revealed the presence of 48 conserved amino acids. On the basis of the alignment, OspC could be divided into an amino-terminal relatively conserved region and a relatively variable region in the central portion. The distance tree obtained divided the ospC sequences into three groups. The first group contained ospC alleles from all (n = 13) sensu stricto strains, the second group contained ospC alleles from seven
 - ***Borrelia*** afzelii strains, and the third group contained ospC alleles from five B. afzelii and all (n = 9) ***Borrelia*** garinii strains. The ratio of the mean number of synonymous (d-S) and nonsynonymous (d-N) nucleotide substitutions per site calculated for B. burgdorferi sensu stricto, B. garinii, and B. afzelii ospC alleles suggested that the polymorphism of OspC is due to positive selection favoring diversity at the amino acid level in the relatively variable region. On the basis of the comparison of 16S rRNA gene sequences,
 - ***Borrelia*** hermsii is more closely related to B. afzelii than to B. burgdorferi sensu stricto and B. garinii. In contrast, the phylogenetic tree obtained for the B. hermsii variable major protein, ***Vmp33***, and 18 OspC amino acid sequences suggested that ***Vmp33*** and OspC from B. burgdorferi sensu stricto strains share a common evolutionary origin.
- TI Evolution of the ***Borrelia*** burgdorferi outer surface protein OspC.
- AB The genes coding for outer surface protein OspC from 22 ***Borrelia*** burgdorferi strains isolated from patients with Lyme ***borreliosis*** were cloned and sequenced. For reference purposes, the 16S rRNA genes from 17 of these strains were sequenced after being. . . group contained ospC alleles from all (n = 13) sensu stricto strains, the second group contained ospC alleles from seven ***Borrelia*** afzelii strains, and the third group contained ospC alleles from five B. afzelii and all (n = 9) ***Borrelia*** garinii strains. The ratio of the mean number of synonymous (d-S) and nonsynonymous (d-N) nucleotide substitutions per site calculated for. . . the amino acid level in the relatively variable region. On the basis of the comparison of 16S rRNA gene sequences, ***Borrelia*** hermsii is more closely related to B.

afzelii than to B. burgdorferi sensu stricto and B. garinii. In contrast, the phylogenetic tree obtained for the B. hermsii variable major protein, ***Vmp33*** , and 18 OspC amino acid sequences suggested that ***Vmp33*** and OspC from B. burgdorferi sensu stricto strains share a common evolutionary origin.

ORGN Classifier

Spirochaetaceae 06112

Super Taxa

Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name

Borrelia afzelii

Borrelia burgdorferi

Taxa Notes

Bacteria, Eubacteria, Microorganisms

- L4 ANSWER 62 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN DUPLICATE 37
- AN 1994:345519 BIOSIS <<LOGINID::20090609>>
- DN PREV199497358519
- TI A family of surface-exposed proteins of 20 kilodaltons in the genus ${\rm ***Borrelia***}$.
- AU Carter, Carol J.; Bergstrom, Sven; Norris, Steven J.; Barbour, Alan G. [Reprint author]
- CS Dep. Microbiol. and Med., Univ. Texas Health Sci. Cent., San Antonio, TX 78284-7758, USA
- SO Infection and Immunity, (1994) Vol. 62, No. 7, pp. 2792-2799. CODEN: INFIBR. ISSN: 0019-9567.
- DT Article
- LA English
- OS Genbank-L24911
 - Entered STN: 8 Aug 1994
 Last Updated on STN: 1 Sep 1994
- Relapsing fever and Lyme disease spirochetes of the genus ***Borrelia*** AΒ display at their surfaces abundant lipoproteins: ***Vmp*** proteins in ***Borrelia*** hermsii and Osp proteins in ***Borrelia*** burgdorferi. ***Vmp*** and Osp proteins largely determine serotype specificity, and neutralizing antibodies of infected or immunized animals are directed at them. For the present study, we examined B. hermsii serotype 33, which is unique among strain HS1 serotypes in the low frequency of switches to other serotypes during infections and in vitro cultivation. Failing to clone the complete ***vmp33*** gene, we accomplished its further characterization by (i) determining three partial amino acid sequences, (ii) designing oligonucleotide primers based on these amino acid sequences, (iii) cloning and sequencing the central portion of ***vmp33*** , and (iv) using outwardly directed primers and the inverse PCR to clone the 5' and 3' ends of the gene and flanking regions. The transcriptional start site was identified by primer extension analysis. ***Vmp33*** was a polypeptide of 211 amino acids; the three partial amino acid sequences were identified in the open reading frame. ***Vmp33*** was found to be more similar to other 20-kDa ***Vmp*** proteins of B. hermsii and to OspC proteins of B. burgdorferi than t was to 35- to 39-kDa ***Vmp*** proteins of the same strain. Moreover, OspC proteins were more similar to ***Vmp33*** than they were to OspA, -B, or -D proteins of B. burgdorferi. These sequence similarities were consistent with Western blot (immunoblot) findings of crossreactions between ***Vmp33*** and OspC with anti- ***Vmp33***

and anti-OspC sera. The promoter for the expressed ***vmp33*** gene

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was found to be different from the expression site for other active
      ***vmp*** genes characterized to date. These results indicate that
      ***Vmp33*** and other small ***Vmp*** 's belong with OspC to a
     genus-wide family of 20-kDa proteins and that expression of these proteins
    may be coordinated with expression of other ***Vmp*** and Osp proteins
     in ***Borrelia*** spp.
    A family of surface-exposed proteins of 20 kilodaltons in the genus
      ***Borrelia***
    Relapsing fever and Lyme disease spirochetes of the genus ***Borrelia***
    display at their surfaces abundant lipoproteins: ***Vmp*** proteins in
      ***Borrelia*** hermsii and Osp proteins in
                                                    ***Borrelia***
     burgdorferi. ***Vmp*** and Osp proteins largely determine serotype
     specificity, and neutralizing antibodies of infected or immunized animals
     are directed at them. For. . . in the low frequency of switches to
     other serotypes during infections and in vitro cultivation. Failing to
     clone the complete ***vmp33*** gene, we accomplished its further
     characterization by (i) determining three partial amino acid sequences,
     (ii) designing oligonucleotide primers based on these amino acid
     sequences, (iii) cloning and sequencing the central portion of
      ***vmp33*** , and (iv) using outwardly directed primers and the inverse
     PCR to clone the 5' and 3' ends of the gene and flanking regions. The
     transcriptional start site was identified by primer extension analysis.
      ***Vmp33*** was a polypeptide of 211 amino acids; the three partial
     amino acid sequences were identified in the open reading frame.
      ***Vmp33*** was found to be more similar to other 20-kDa
     proteins of B. hermsii and to OspC proteins of B. burgdorferi than t was
     to 35- to 39-kDa ***Vmp*** proteins of the same strain. Moreover,
    OspC proteins were more similar to ***Vmp33*** than they were to OspA,
    -B, or -D proteins of B. burgdorferi. These sequence similarities were
     consistent with Western blot (immunoblot) findings of crossreactions
             ***Vmp33*** and OspC with anti- ***Vmp33*** and anti-OspC
     sera. The promoter for the expressed ***vmp33*** gene was found to be
     different from the expression site for other active ***vmp*** genes
     characterized to date. These results indicate that ***Vmp33***
     other small ***Vmp*** 's belong with OspC to a genus-wide family of
     20-kDa proteins and that expression of these proteins may be coordinated
    with expression of other ***Vmp*** and Osp proteins in
      ***Borrelia***
                      spp.
       sequence data; nucleotide sequence; L24911: Genbank
    Miscellaneous Descriptors
       CLONING STRATEGY; HOMOLOGY; METHOD; OSPC PROTEIN; PROMOTER ANALYSIS;
       TRANSCRIPTION START SITE; ***VMP33***
                                               GENE
ORGN Classifier
       Spirochaetaceae 06112
     Super Taxa
       Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms
     Organism Name
           ***Borrelia***
                          burgdorferi
           ***Borrelia*** hermsii
     Taxa Notes
       Bacteria, Eubacteria, Microorganisms
    ANSWER 63 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on
                                                      DUPLICATE 38
    1994:498441 BIOSIS <<LOGINID::20090609>>
    PREV199497511441
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ΤI

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ΑN DN

- TI Antigen diversity in the bacterium B. hermsii through "somatic' mutations in rearrange ***vmp*** genes.
- AU Restrepo, Blanca I.; Barbour, Alan G.
- CS Dep. Microbiol., Univ. Texas Health Sci. Cent., San Antonio, TX 78284-7758, USA
- SO Cell, (1994) Vol. 78, No. 5, pp. 867-876. CODEN: CELLB5. ISSN: 0092-8674.
- DT Article
- LA English
- ED Entered STN: 28 Nov 1994
 Last Updated on STN: 28 Nov 1994
- AB B. hermsii counters host immunity with multiphasic antigenic variation. This is conferred by interplasmidic and intraplasmidic rearrangements of ***vmp*** genes. In several Independent events, activation of a silent ***vmp*** gene through intraplasmidic deletions but not interplasmidic recombinations was followed by the appearance at its 5' end of multiple mutations that were not present in the silent gene. The prevalence of mutant alleles in postswitch populations increased during infections. Differences between the silent and expressed genes were at the same nucleotides at which ***vmp*** pseudogenes differed, suggesting these were templates for postswitch gene conversions. The mechanism of this bacterium to generate diversity, namely, intramolecular deletions followed by mutations in the rearranged gene, mirrors the strategy used by vertebrate hosts to eliminate it.
- TI Antigen diversity in the bacterium B. hermsii through "somatic' mutations in rearrange ***vmp*** genes.
- AB B. hermsii counters host immunity with multiphasic antigenic variation.

 This is conferred by interplasmidic and intraplasmidic rearrangements of

 vmp genes. In several Independent events, activation of a silent

 vmp gene through intraplasmidic deletions but not interplasmidic

 recombinations was followed by the appearance at its 5' end of multiple

 mutations. . . in postswitch populations increased during infections.

 Differences between the silent and expressed genes were at the same

 nucleotides at which ***vmp*** pseudogenes differed, suggesting these

 were templates for postswitch gene conversions. The mechanism of this

 bacterium to generate diversity, namely, intramolecular. . .
- IT Miscellaneous Descriptors

BACTERIAL VIRULENCE; HOST IMMUNITY; INTRAMOLECULAR DELETION; POSTSWITCH GENE CONVERSION; PSI- ***VMP26*** PSEUDOGENE; SILENT SITE

ORGN . .

Mammals, Nonhuman Vertebrates, Nonhuman Mammals, Rodents, Vertebrates ORGN Classifier

Spirochaetaceae 06112

Super Taxa

Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name

Borrelia hermsii

Taxa Notes

Bacteria, Eubacteria, Microorganisms

- L4 ANSWER 64 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN DUPLICATE 39
- AN 1994:128747 BIOSIS <<LOGINID::20090609>>
- DN PREV199497141747
- TI Variability of a bacterial surface protein and disease expression in a possible mouse model of systemic lyme ***borreliosis*** .
- AU Cadavid, Diego; Thomas, D. Denee; Crawley, Ronald; Barbour, Alan G.

[Reprint author]

- CS Dep. Microbiol., Univ. Texas Health Sci. Center, 7703 Floyd Curl Dr., San Antonio, TX 78284-7758, USA
- SO Journal of Experimental Medicine, (1994) Vol. 179, No. 2, pp. 631-642. CODEN: JEMEAV. ISSN: 0022-1007.
- DT Article
- LA English
- ED Entered STN: 24 Mar 1994 Last Updated on STN: 24 Mar 1994
- AΒ During persistent infection of scid mice with ***Borrelia*** turicatae, an agent of relapsing fever and neuroborreliosis, there was variation in the surface proteins the bacteria expressed and in disease manifestations over time. Two serotypes, A and B, were isolated from the mice, cloned by limiting dilution, and further characterized. The only discernible difference between the two variants was in the size of the major surface protein they expressed: serotype A had a variable major protein (***Vmp***) of 23,000, and serotype B had a ***Vmp*** of 20,000. When other scid mice were inoculated with clonal populations of A and B, the infections were similar with respect to onset and degree of spirochetemia, involvement of the eye and heart, and occurrence of a peripheral vestibular disorder. However, there were differences between the serotypes in other respects: (a) serotype B but not A caused reddened and significantly enlarged joints, markedly impaired performance on a walking bar, and severe arthritis by histologic examination; (b) serotype A but not B invaded the central nervous system during early infection; and (c) serotype A penetrated monolayers of human umbilical vein endothelial cells more readily than did serotype B. The combination of arthritis, myocarditis, and neurologic disease resembled human Lyme

borreliosis . The findings indicate that differences in disease expression are determined by variable surface proteins of the bacterium and that scid mouse infections with B. turicatae provide a model for the study of the pathogenesis of Lyme ***borreliosis*** and other persistent spirochetal diseases.

- TI Variability of a bacterial surface protein and disease expression in a possible mouse model of systemic lyme ***borreliosis*** .
- During persistent infection of scid mice with ***Borrelia*** AΒ turicatae, an agent of relapsing fever and neuroborreliosis, there was variation in the surface proteins the bacteria expressed and in. . . two variants was in the size of the major surface protein they expressed: serotype A had a variable major protein (***Vmp***) of 23,000, and serotype B had a ***Vmp*** of 20,000. When other scid mice were inoculated with clonal populations of A and B, the infections were similar with. . . vein endothelial cells more readily than did serotype B. The combination of arthritis, myocarditis, and neurologic disease resembled ***borreliosis*** . The findings indicate that differences human Lyme in disease expression are determined by variable surface proteins of the bacterium and that scid mouse infections with B. turicatae provide a model for the study of the pathogenesis of Lyme ***borreliosis*** and other persistent spirochetal diseases.

ORGN . .

Mammals, Nonhuman Vertebrates, Nonhuman Mammals, Rodents, Vertebrates ORGN Classifier

Spirochaetaceae 06112

Super Taxa

Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name

Borrelia turicatae

Taxa Notes

Bacteria, Eubacteria, Microorganisms

- L4 ANSWER 65 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN DUPLICATE 40
- AN 1994:435145 BIOSIS <<LOGINID::20090609>>
- DN PREV199497448145
- TI Activation of a ***vmp*** pseudogene in ***Borrelia*** hermsii: An alternate mechanism of antigenic variation during relapsing fever.
- AU Restrepo, B. I.; Carter, C. J.; Barbour, A. G. [Reprint author]
- CS Dep. Med., Univ. Tex. Health Sci. Cent., San Antonio, TX 78284-7758, USA
- SO Molecular Microbiology, (1994) Vol. 13, No. 2, pp. 287-299. CODEN: MOMIEE. ISSN: 0950-382X.
- DT Article
- LA English
- ED Entered STN: 11 Oct 1994

 Last Updated on STN: 11 Oct 1994
- AB The relapsing fever agent, ***Borrelia*** hermsii, undergoes multiphasic antigenic variation to evade its host's immune response. A frequently observed switch is serotype 7 to 26. Unlike silent ***vmp*** genes previously characterized, the transcriptionally silent ***vmp26*** sequence was a pseudogene in lacking a start codon. In serotype 7 the location of the silent ***vmp26*** sequence just downstream of ***vmp7*** on the expression plasmid, as well as on the silent plasmid, was also unique. The demonstration of a predicted circular recombination product in serotype 7 but not serotype 21 populations indicates that the pseudogene was activated by an intramolecular recombination producing a deletion of DNA between 20-nucleotide direct repeats in ***vmp7*** and PSI ***vmp26***.
- TI Activation of a ***vmp*** pseudogene in ***Borrelia*** hermsii: An alternate mechanism of antigenic variation during relapsing fever.
- AB The relapsing fever agent, ***Borrelia*** hermsii, undergoes multiphasic antigenic variation to evade its host's immune response. A frequently observed switch is serotype 7 to 26. Unlike silent ***vmp*** genes previously characterized, the transcriptionally silent ***vmp26*** sequence was a pseudogene in lacking a start codon. In serotype 7 the location of the silent ***vmp26*** sequence just downstream of ***vmp7*** on the expression plasmid, as well as on the silent plasmid, was also unique. The demonstration of a predicted circular. . . indicates that the pseudogene was activated by an intramolecular recombination producing a deletion of DNA between 20-nucleotide direct repeats in ***vmp7*** and PSI ***vmp26***.

ORGN . . .

Notes

Animals, Chordates, Humans, Mammals, Primates, Vertebrates ORGN Classifier

Spirochaetaceae 06112

Super Taxa

Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name

Borrelia hermsii

Taxa Notes

Bacteria, Eubacteria, Microorganisms

- L4 ANSWER 66 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN DUPLICATE 41
- AN 1994:357938 BIOSIS <<LOGINID::20090609>>

- DN PREV199497370938
- TI Homology between ***Borrelia*** burgdorferi OspC and members of the family of ***Borrelia*** hermsii variable major proteins.
- AU Margolis, Neil; Hogan, Daniel; Cieplak., Witold, Jr.; Schwan, Tom G.; Rosa, Patricia A. [Reprint author]
- CS Lab. Microbial Structure and Function, Rocky Mountain Lab., Natl. Inst. Health, Natl. Inst. Allergy and Infectious Dis., Hamilton, MT 59840, USA
- SO Gene (Amsterdam), (1994) Vol. 143, No. 1, pp. 105-110. CODEN: GENED6. ISSN: 0378-1119.
- DT Article
- LA English
- ED Entered STN: 23 Aug 1994 Last Updated on STN: 23 Aug 1994
- AΒ Synthesis of the ***Borrelia*** burgdorferi outer surface protein C (OspC) is quite variable. We have cloned and sequenced the ospC gene from B. burgdorferi isolate CA-11.2A, a clone in which ospC expression varies. The 5' flanking region of the gene contains at least two consensus promoter regions, as well as two large overlapping inverted repeats. Sequence comparison to other OspC proteins indicated that the CA-11.2A OspC is as closely related to OspC from two different genospecies of Lyme disease spirochetes as it is to OspC from the prototype B. burgdorferi strain, B31. Comparisons of the OspC amino acid (aa) sequence with those in aa sequence databases revealed partial identity with the variable major proteins ***Vmp3*** and ***Vmp24*** of B. hermsii, a causative agent of tick-borne relapsing fever. An ospC probe hybridized to B. hermsii restriction fragments and linear plasmids that also were recognized by the ***vmp3*** and ***vmp24*** probes. OspC and ***Vmp*** appear to be related, but their synthesis is regulated these differently in the two species of spirochetes. This represents a fascinating example of the evolution of the number, position, regulation and perhaps function of homologous genes in two related pathogens. These parameters may relate to characteristic properties of the pathogens and their separate tick vectors.
- TI Homology between ***Borrelia*** burgdorferi OspC and members of the family of ***Borrelia*** hermsii variable major proteins.
- AB Synthesis of the ***Borrelia*** burgdorferi outer surface protein C (OspC) is quite variable. We have cloned and sequenced the ospC gene from B. burgdorferi. . . the OspC amino acid (aa) sequence with those in aa sequence databases revealed partial identity with the variable major proteins ***Vmp3*** and ***Vmp24*** of B. hermsii, a causative agent of tick-borne relapsing fever. An ospC probe hybridized to B. hermsii restriction fragments and linear plasmids that also were recognized by the ***vmp3*** and ***vmp24*** probes. OspC and these ***Vmp*** appear to be related, but their synthesis is regulated differently in the two species of spirochetes. This represents a fascinating. . .

ORGN Classifier

Spirochaetaceae 06112

Super Taxa

Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name

Borrelia burgdorferi ***Borrelia*** hermsii

Taxa Notes

Bacteria, Eubacteria, Microorganisms

L4 ANSWER 67 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on

STN DUPLICATE 42

- AN 1993:299705 BIOSIS <<LOGINID::20090609>>
- DN PREV199396017930
- TI Intragenic recombination and a chimeric outer membrane protein in the relapsing fever agent ***Borrelia*** hermsii.
- AU Kitten, Todd; Barrera, Adrian V.; Barbour, Alan G. [Reprint author]
- CS Dep. Microbiol. Med., Univ. Texas Health Sci. Center, San Antonio, Texas 78284-7758, USA
- SO Journal of Bacteriology, (1993) Vol. 175, No. 9, pp. 2516-2522. CODEN: JOBAAY. ISSN: 0021-9193.
- DT Article
- LA English
- ED Entered STN: 23 Jun 1993 Last Updated on STN: 23 Jun 1993
- AB The spirochete ***Borrelia*** hermsii, a relapsing fever agent, evades the host's immune response through multiphasic antigenic variation. Antigen switching results from sequential expression of genes for serotype-specific outer membrane proteins known as variable major proteins (***Vmp*** 's); of the 25 serotypes that have been identified for the HS1 strain, serotypes 7 and 21 have been studied in greatest detail. In the present study, an atypical variant was predominant in the relapse from a serotype 21 infection in mice; relapse cells were bound by monoclonal antibodies specific for ***Vmp21*** as well as antibodies specific for ***Vmp7***. In Western blots (immunoblots), the variant had a single ***Vmp*** that was reactive with monoclonal antibodies representing

both

serotypes. The gene encoding this ***Vmp*** , ***vmp7*** /21, was cloned and characterized by restriction mapping and sequence analysis to determine the likely recombination event. Whereas the 5' end of ***vmp7*** /21 was identical to that of ***vmp21*** , its 3' end and flanking sequences were identical to the 3' end of ***vmp7*** . Unlike other ***vmp*** genes examined thus far, the ***vmp7*** /21 gene existed only in an expressed form; a silent, storage form of the gene was not detected. We conclude that the ***vmp7*** /21 gene was created by an intragenic recombination between the formerly expressed ***vmp21*** gene and a silent ***vmp7*** gene. This finding suggests that the lack of cross-reactivity between variants, which is usually observed, results from immunoselection against variants possessing chimeric ***Vmp*** 's rather than from a switching mechanism that excludes

gene replacements.

- TI Intragenic recombination and a chimeric outer membrane protein in the relapsing fever agent ***Borrelia*** hermsii.
- AB The spirochete ***Borrelia*** hermsii, a relapsing fever agent, evades the host's immune response through multiphasic antigenic variation. Antigen switching results from sequential expression of genes for serotype-specific outer membrane proteins known as variable major proteins (***Vmp*** 's); of the 25 serotypes that have been identified for the HS1 strain, serotypes 7 and 21 have been studied in. . . predominant in the relapse from a serotype 21 infection in mice; relapse cells were bound by monoclonal antibodies specific for ***Vmp21*** as well as antibodies specific for ***Vmp7***. In Western blots (immunoblots), the variant had a single ***Vmp*** that was reactive with monoclonal antibodies representing both serotypes. The gene encoding this ***Vmp*** , ***vmp7*** /21, was cloned and characterized by restriction mapping and sequence analysis to determine the likely recombination event. Whereas the 5' end of ***vmp7*** /21 was

identical to that of $\ \ ^{***}vmp21^{***}$, its 3' end and flanking sequences were identical to the 3' end of ***vmp7*** . Unlike other ***vmp*** genes examined thus far, the ***vmp7*** /21 gene existed only in an expressed form; a silent, storage form of the gene was not detected. We conclude that the ***vmp7*** /21 gene was created by an intragenic recombination between the formerly expressed ***vmp21*** gene and a ***vmp7*** gene. This finding suggests that the lack of cross-reactivity between variants, which is usually observed, results from immunoselection against variants possessing chimeric ***Vmp*** 's rather than from a switching mechanism that excludes partial gene replacements. Miscellaneous Descriptors ANTIGEN SWITCHING MECHANISM; GENE MAPPING; IMMUNE SELECTION; RESTRICTION MAPPING; ***VMP21*** GENE; ***VMP7*** GENE; ***VMP7*** -21 GENE ORGN Classifier Spirochaetaceae 06112 Super Taxa Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name ***Borrelia*** hermsii Taxa Notes Bacteria, Eubacteria, Microorganisms ANSWER 68 OF 87 MEDLINE on STN 1994184809 MEDLINE <<LOGINID::20090609>> PubMed ID: 8137122 Linear DNA of ***Borrelia*** species and antigenic variation. Barbour A G Dept of Microbiology, University of Texas Health Science Center, San Antonio 78284. AI24424 (United States NIAID NIH HHS) AI29731 (United States NIAID NIH HHS) AR41507 (United States NIAMS NIH HHS) Trends in microbiology, (1993 Sep) Vol. 1, No. 6, pp. 236-9. Journal code: 9310916. ISSN: 0966-842X. ENGLAND: United Kingdom Journal; Article; (JOURNAL ARTICLE) (RESEARCH SUPPORT, U.S. GOV'T, NON-P.H.S.) (RESEARCH SUPPORT, U.S. GOV'T, P.H.S.) English Priority Journals 199404 Entered STN: 9 May 1994 Last Updated on STN: 9 May 1994 Entered Medline: 28 Apr 1994 Members of the genus ***Borrelia*** may be unique among prokaryotic organisms in having a polyploid genome that is mostly linear. The smaller linear duplex replicons in these organisms have been called plasmids, but there is justification for designating them minichromosomes instead. The antigenic identities of the agents of Lyme disease and relapsing fever are largely determined by these extrachromosomal genes.

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LΑ

FS EM

- ΤI Linear DNA of ***Borrelia*** species and antigenic variation.
- Members of the genus ***Borrelia*** may be unique among prokaryotic organisms in having a polyploid genome that is mostly linear. The smaller linear duplex replicons. . .

```
СТ
     *Antigenic Variation: GE, genetics
     Bacterial Outer Membrane Proteins: GE, genetics
     Base Sequence
         ****Borrelia: GE, genetics***
         *** Borrelia: IM, immunology***
     *DNA, Bacterial: GE, genetics
     Genome, Bacterial
     Molecular Sequence Data
     Plasmids: GE, genetics
GEN osp; ***vmp***
L4
    ANSWER 69 OF 87 CAPLUS COPYRIGHT 2009 ACS on STN DUPLICATE 43
AN
    1994:74507 CAPLUS <<LOGINID::20090609>>
    120:74507
OREF 120:13359a,13362a
    Experimental infection of the mouse brain by a relapsing fever
TΤ
       ***Borrelia*** species: a molecular analysis
ΑU
    Cadavid, Diego; Bundoc, Virgilio; Barbour, Alan G.
CS
    Health Sci. Cent., Univ. Texas, San Antonio, TX, 78284-7758, USA
SO
    Journal of Infectious Diseases (1993), 168(1), 143-51
    CODEN: JIDIAO; ISSN: 0022-1899
DT
    Journal
    English
LA
    The spirochetal disease relapsing fever is notable not only for
AB
    multiphasic antigenic variation but also for central neurol.
    manifestations. To further characterize involvement of the brain in this
    disorder, immunocompetent and immunodeficient mice were infected with
       ***Borrelia*** hermsii. Immunodeficient mice were treated while
     spirochetemic with neutralizing IgM monoclonal antibodies to the infecting
     serotype. Blood, cerebrospinal fluid, and brain tissue were examd. by
     culture and polymerase chain reaction. In immunocompetent mice, antigenic
     variation occurred in the brain as well as in the blood. In
     immunodeficient mice, the infecting serotype was still present in the
    brain after it had been eliminated from the blood by the administered
    antibodies. These latter results cannot be accounted for by contamination
    of brain tissue and cerebrospinal fluid by blood and, hence, establish the
    direct involvement of the central nervous system in this exptl. infection.
    Experimental infection of the mouse brain by a relapsing fever
ΤI
      ***Borrelia*** species: a molecular analysis
     . . . central neurol. manifestations. To further characterize
AB
     involvement of the brain in this disorder, immunocompetent and
     immunodeficient mice were infected with ***Borrelia*** hermsii.
     Immunodeficient mice were treated while spirochetemic with neutralizing
     IgM monoclonal antibodies to the infecting serotype. Blood, cerebrospinal
     fluid, and. . .
      ***Borrelia*** brain antigen ***Vmp***
ST
                                                  protein sequence; gene
       ***vmp***
                   ***Borrelia*** brain sequence
ΙT
    Antigens
    RL: BIOL (Biological study)
        ( ***Borrelia*** hermsii variable major protein New ***Vmp***
       as, in relapsing fever)
ΙT
    Gene, microbial
    RL: BIOL (Biological study)
        (new ***vmp*** , nucleotide sequence of, of ***Borrelia***
       hermsii, relapsing fever in relation to)
ΙT
    Deoxyribonucleic acid sequences
        (of gene new ***vmp*** of ***Borrelia*** hermsii, relapsing
```

```
fever in relation to)
ΙT
    Protein sequences
        (of variable major protein New ***Vmp*** of ***Borrelia***
       hermsii, relapsing fever in relation to)
ΙT
       ***Borrelia*** hermsii
        (variable major protein New ***Vmp*** of, amino acid sequence of,
       relapsing fever in relation to)
ΤТ
    Brain, disease
        (infection, with
                          ***Borrelia*** hermsii, variable major protein New
         ***Vmp*** in)
ΙT
    Fever and Hyperthermia
        (relapsing, from
                          ***Borrelia*** hermsii infection of brain,
       variable major protein New ***Vmp***
                                               in)
ΙT
     152522-18-4, Protein ( ***Borrelia*** hermsii clone bp7E variable major
     protein New ***Vmp*** )
    RL: PRP (Properties)
        (amino acid sequence of)
ΙT
    152522-17-3, DNA ( ***Borrelia***
                                        hermsii clone bp7E gene new
      ***vmp*** mRNA-complementary)
     RL: PRP (Properties)
        (nucleotide sequence of)
    ANSWER 70 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on
L4
                                                       DUPLICATE 44
    1993:94463 BIOSIS <<LOGINID::20090609>>
ΑN
    PREV199395049659
DN
    Subtelomeric expression regions of
                                        ***Borrelia*** hermsii linear
    plasmids are highly polymorphic.
ΑU
    Restrepo, B. I.; Kitten, T.; Carter, C. J.; Infante, D.; Barbour, A. G.
     [Reprint author]
CS
     Dep. Microbiol., Univ. Tex. Health Sci. Cent., San Antonio, Tex.
    78284-7758, USA
    Molecular Microbiology, (1992) Vol. 6, No. 22, pp. 3299-3311.
SO
    CODEN: MOMIEE. ISSN: 0950-382X.
DT
    Article
LA
    English
    DDBJ-LO4786; DDBJ-LO4787; DDBJ-LO4788; DDBJ-LO4789; DDBJ-MS7256;
OS
     DDBJ-Z11876; EMBL-LO4786; EMBL-LO4787; EMBL-LO4788; EMBL-LO4789;
     EMBL-MS7256; EMBL-Z11876; Genbank-LO4786; Genbank-LO4787; Genbank-LO4788;
    Genbank-LO4789; Genbank-MS7256; Genbank-Z11876
ED
    Entered STN: 9 Feb 1993
    Last Updated on STN: 17 Apr 1993
      ***Borrelia*** hermsii, a relapsing fever agent, undergoes multiphasic
AΒ
     antigenic variation to evade its host's immune response. Serotype
     specificity is determined by variable membrane lipoproteins,
                                                                  ***Vmps***
     , which are expressed from genes located near the end of a linear plasmid.
     Using the polymerase chain reaction and primers representing the promoter
     of the active ***vmp*** and a conserved telomeric sequence, we
     characterized the subtelomeric expression regions of the 25 known
     serotypes of strain HS1. The distance from the promoter to the telomere
     fell into three size classes of approximately 1.0, 1.5, and 2.5 kilobases.
     In the sequenced serotypes the size differences were accounted for by
    variable lengths of the ***vmp*** genes and intervening sequences
    between 3' end of the ***vmp*** gene and the start of a downstream
    homology block. The degree of nucleotide identify between different
```

vmp genes, or between the different 3' flanking DNA varied from 39-78%. Thus, there is length and sequence variability not only between

vmp genes themselves but also between the 3' flanking regions of ***vmp*** genes.

TI Subtelomeric expression regions of ***Borrelia*** hermsii linear plasmids are highly polymorphic.

Borrelia hermsii, a relapsing fever agent, undergoes multiphasic AB antiquenic variation to evade its host's immune response. Serotype specificity is determined by variable membrane lipoproteins, ***Vmps*** , which are expressed from genes located near the end of a linear plasmid. Using the polymerase chain reaction and primers representing the promoter of the active ***vmp*** and a conserved telomeric sequence, we characterized the subtelomeric expression regions of the 25 known serotypes of strain HS1. The. . . 1.0, 1.5, and 2.5 kilobases. In the sequenced serotypes the size differences were accounted for by variable lengths of the ***vmp*** genes and intervening sequences between 3' end of the ***vmp*** gene and the start of a downstream homology block. The degree of nucleotide identify between different ***vmp*** genes, or between the different 3' flanking DNA varied from 39-78%. Thus, there is length and sequence variability not only between ***vmp*** genes themselves but also between the 3' flanking regions of ***vmp*** genes.

ORGN Classifier

Spirochaetaceae 06112

Super Taxa

Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name

Borrelia hermsii

Taxa Notes

Bacteria, Eubacteria, Microorganisms

- L4 ANSWER 71 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN DUPLICATE 45
- AN 1993:3952 BIOSIS <<LOGINID::20090609>>
- DN PREV199395003952
- TI The relapsing fever agent ***Borrelia*** hermsii has multiple copies of its chromosome and linear plasmids.
- AU Kitten, Todd [Reprint author]; Barbour, Alan G.
- CS Dep. Microbiol., University Texas Health Sci. Center, San Antonio, Texas 78284, USA
- SO Genetics, (1992) Vol. 132, No. 2, pp. 311-324. CODEN: GENTAE. ISSN: 0016-6731.
- DT Article
- LA English
- ED Entered STN: 10 Dec 1992
 - Last Updated on STN: 10 Dec 1992
- ***Borrelia*** hermsii, a spirochete which causes relapsing fever in humans and other mammals, eludes the immune response by antigenic variation of the " ***Vmp*** " proteins. This occurs by replacement of an expressed ***vmp*** gene with a copy of a silent ***vmp*** gene. Silent and expressed ***vmp*** genes are located on separate linear plasmids. To further characterize ***vmp*** recombination, copy numbers were determined for two linear plasmids and for the 1-megabase chromosome by comparing hybridization of probes to native DNA with hybridization to recombinant plasmids containing ***borrelial*** DNA. Plasmid copy numbers were also estimated by ethicium bromide fluorescence. Total cellular DNA content was determined by spectrophotometry. For ***borrelias*** grown in mice, copy numbers and 95% confidence intervals were 14 (12-17) for an expression plasmid, 8

(7-9) for a silent plasmid, and 16 (13-18) for the chromosome. ***Borrelias*** grown in broth medium had one-fourth to one-half this number of plasmids and chromosomes. Staining of cells with 4',6-diamidino-2-phenylindole revealed DNA to be distributed throughout most of the spirochete's length. These findings indicate that ***borrelias*** organize their total cellular DNA into several complete genomes and that cells undergoing serotype switches do one or more of the following: (1) coexpress ***Vmps*** from switched and unswitched expression plasmids for at least three to five generations, (2) suppress transcription from some expression plasmid copies, or (3) partition expression plasmids nonrandomly. The lower copy number of the silent plasmid indicates that nonreciprocal ***Vmp*** gene recombination may result from loss of recombinant silent plasmids by segregation. The relapsing fever agent ***Borrelia*** hermsii has multiple copies of its chromosome and linear plasmids. ***Borrelia*** hermsii, a spirochete which causes relapsing fever in humans and other mammals, eludes the immune response by antigenic variation of the " ***Vmp*** " proteins. This occurs by replacement of an expressed ***vmp*** gene with a copy of a silent ***vmp*** gene. Silent and expressed ***vmp*** genes are located on separate linear plasmids. To further characterize ***vmp*** recombination, copy numbers were determined for two linear plasmids and for the 1-megabase chromosome by comparing hybridization of probes to native DNA with hybridization to recombinant plasmids containing ***borrelial*** DNA. Plasmid copy numbers were also estimated by ethidium bromide fluorescence. Total cellular DNA content was determined by spectrophotometry. For ***borrelias*** grown in mice, copy numbers and 95% confidence intervals were 14 (12-17) for an expression plasmid, 8(7-9) for a silent plasmid, and 16 (13-18) for the chromosome. ***Borrelias*** grown in broth medium had one-fourth to one-half this number of plasmids and chromosomes. Staining of cells with 4',6-diamidino-2-phenylindole revealed DNA to be distributed throughout most of the spirochete's length. These findings indicate that ***borrelias*** organize their total cellular DNA into several complete genomes and that cells undergoing serotype switches do one or more of the following: (1) coexpress ***Vmps*** from switched and unswitched expression plasmids for at least three to five generations, (2) suppress transcription from some expression plasmid copies, or (3) partition expression plasmids nonrandomly. The lower copy number of the silent plasmid indicates that nonreciprocal ***Vmp*** gene recombination may result from loss of recombinant silent plasmids by segregation. Miscellaneous Descriptors ***VMP*** GENE EXPRESSION; ***VMP*** ANTIGENIC VARIATION; PROTEIN ORGN Classifier Spirochaetaceae 06112 Spirochaetales; Spirochetes; Eubacteria; Bacteria; Microorganisms Organism Name ***Borrelia*** hermsii Taxa Notes Bacteria, Eubacteria, Microorganisms ANSWER 72 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on DUPLICATE 46 1991:510076 BIOSIS <<LOGINID::20090609>>

ΤI

AB

ΤT

T.4

AN

DN

PREV199141110791; BR41:110791

- ANTIGENIC VARIATION IN ***BORRELIA*** ΤТ GIRONS I S [Reprint author]; BARBOUR A G AU UNITE LEPTOSPIRES, INST PASTEUR, 75724 PARIS CEDEX 15 CS Research in Microbiology, (1991) Vol. 142, No. 6, pp. 711-718. CODEN: RMCREW. ISSN: 0923-2508. Article DΤ FS T.A ENGLISH ED Entered STN: 14 Nov 1991 Last Updated on STN: 14 Nov 1991 ΤI ANTIGENIC VARIATION IN ***BORRELIA*** . ΙT Miscellaneous Descriptors REVIEW ***BORRELIA*** -HERMSII RELAPSING FEVER MAJOR SURFACE PROTEIN ***VMP*** GENES PLASMID GENE TRANSPOSITION GENE ACTIVATION TELOMERIC SITE REARRANGEMENT L4ANSWER 73 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on DUPLICATE 47 1991:226831 BIOSIS <<LOGINID::20090609>> ΑN DN PREV199191118291; BA91:118291 VARIABLE ANTIGEN GENES OF THE RELAPSING FEVER AGENT ***BORRELIA*** ΤI -HERMSII ARE ACTIVATED BY PROMOTER ADDITION. BARBOUR A G [Reprint author]; BURMAN N; CARTER C J; KITTEN T; BERGSTROM S ΑU CS DEP MICROBIOL AND MED, UNIV TEX HEALTH SCI CENT, SAN ANTONIO, TEX 78284, Molecular Microbiology, (1991) Vol. 5, No. 2, pp. 489-494. SO CODEN: MOMIEE. ISSN: 0950-382X. Article DT FS ENGLISH LA ED Entered STN: 9 May 1991 Last Updated on STN: 9 May 1991 ***Borrelia*** hermsii, an agent of relapsing fever, avoids the host's AB immune response by means of multiphasia antigenic variation. Serotype specificity is determined by variable antigens called the ***Vmp*** lipoproteins. Through recombination between linear plasmids a formerly silent ***vmp*** gene replaces another ***vmp*** gene at a telomeric expression locus. We examined strain HS1 ***borreliae*** before and after a switch from serotype 7 to serotype 21. The nucleotide sequences of 5' regions of silent and expressed ***vmp7*** and ***vmp21*** were determined. Silent and active ***vmp7*** ***vmp21*** genes shared a block of homologous sequences surrounding their 5' ends. Sequences upstream of silent ***vmp7*** and ${\rm ***vmp21***}$ genes lacked the promoter and substantially differed from each other. In this antigenic switch a ***vmp*** gene was activated by a recombination that placed it downstream of a promoter. VARIABLE ANTIGEN GENES OF THE RELAPSING FEVER AGENT ***BORRELIA*** TΤ -HERMSII ARE ACTIVATED BY PROMOTER ADDITION.
- ***Borrelia*** hermsii, an agent of relapsing fever, avoids the host's immune response by means of multiphasia antigenic variation. Serotype specificity is determined by variable antigens called the ***Vmp*** lipoproteins. Through recombination between linear plasmids a formerly silent ***vmp*** gene replaces another ***vmp*** gene at a telomeric expression locus. We examined strain HS1 ***borreliae*** before and after a switch from serotype 7 to serotype 21. The nucleotide sequences of 5' regions of silent and expressed ***vmp7*** and ***vmp21*** were determined. Silent and active ***vmp7*** and

- ***vmp21*** genes shared a block of homologous sequences surrounding their 5' ends. Sequences upstream of silent ***vmp7*** and ***vmp21*** genes lacked the promoter and substantially differed from each other. In this antigenic switch a ***vmp*** gene was activated by a recombination that placed it downstream of a promoter.
- IT Miscellaneous Descriptors
 LIPOPROTEIN ***VMP*** GENE RECOMBINATION NUCLEOTIDE SEQUENCE
 MOLECULAR SEQUENCE DATA
- L4 ANSWER 74 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN DUPLICATE 48
- AN 1991:113384 BIOSIS <<LOGINID::20090609>>
- DN PREV199191060774; BA91:60774
- TI TANDEM INSERTION SEQUENCE-LIKE ELEMENTS DEFINE THE EXPRESSION SITE FOR VARIABLE ANTIGEN GENES OF ***BORRELIA*** -HERMSII.
- AU BARBOUR A G [Reprint author]; CARTER C J; BURMAN N; FREITAG C S; GARON C F; BERGSTROM S
- CS DEP MICROBIOL, UNIV TEXAS HEALTH SCI CENT, SAN ANTONIO, TX 78284, USA
- SO Infection and Immunity, (1991) Vol. 59, No. 1, pp. 390-397. CODEN: INFIBR. ISSN: 0019-9567.
- DT Article
- FS BA
- LA ENGLISH
- ED Entered STN: 27 Feb 1991 Last Updated on STN: 27 Feb 1991
- The spirochete ***Borrelia*** hermsii avoids the immune response of AΒ its mammalian host through multiphasic antigenic variation. Serotype specificity is determined by variable antigens, ***Vmp*** proteins, in the outer membrane. Through nonreciprocal recombination between linear plasmids, a formerly silent ***vmp*** gene replaces another gene downstream from a common expression site. To futher characterize this activating site, we determined the nucleotide sequence of 6.9 kb of the common upstream expression region of strain HS1 of B. hermsii. Preceding the ***vmp*** gene promoter and a poly(dT .cntdot. dA) run were three imperfectly repeated segments of 2 kb. Each of the 2-kb segments contained 1-kb elements with inverted repeats of approximately 0.2 kb each at their termini. The potential of the 1-kb elements to form stem-and-loop structures was demonstrated by heteroduplex analysis. There was no evidence of the presence of the elements elsewhere in the genome of B. hermsii. One or more of these elements may confer the unidirectionality that characterizes ***vmp*** gene switches. TANDEM INSERTION SEQUENCE-LIKE ELEMENTS DEFINE THE EXPRESSION SITE FOR ТΤ
- TI TANDEM INSERTION SEQUENCE-LIKE ELEMENTS DEFINE THE EXPRESSION SITE FOR VARIABLE ANTIGEN GENES OF ***BORRELIA*** -HERMSII.
- AB The spirochete ***Borrelia*** hermsii avoids the immune response of its mammalian host through multiphasic antigenic variation. Serotype specificity is determined by variable antigens, ***Vmp*** proteins, in the outer membrane. Through nonreciprocal recombination between linear plasmids, a formerly silent ***vmp*** gene replaces another ***vmp*** gene downstream from a common expression site. To futher characterize this activating site, we determined the nucleotide sequence of 6.9 kb of the common upstream expression region of strain HS1 of B. hermsii. Preceding the ***vmp*** gene promoter and a poly(dT .cntdot. dA) run were three imperfectly repeated segments of 2 kb. Each of the
 - dA) run were three imperfectly repeated segments of 2 kb. Each of the 2-kb. . . elements elsewhere in the genome of B. hermsii. One or more of these elements may confer the unidirectionality that characterizes ***vmp*** gene switches.

- L4 ANSWER 75 OF 87 CABA COPYRIGHT 2009 CABI on STN
- AN 91:89224 CABA <<LOGINID::20090609>>
- DN 19910506221
- TI Antigenic variation in relapsing fever ***Borrelia*** species
- AU Barbour, A. G.; Iglewski, B.H. [EDITOR]; Clark, V.L. [EDITOR]
- CS Departments of Microbiology and Medicine, University of Texas Health Science Center, San Antonio, TX 78284, USA.
- SO The bacteria: a treatise on structure and function. Volume XI. Molecular basis of bacterial pathogenesis, (1990) pp. 155-176. 44 ref. Publisher: Academic Press, Inc. San Diego, California
- CY United States
- DT Miscellaneous
- LA English
- ED Entered STN: 1 Nov 1994

 Last Updated on STN: 1 Nov 1994
- AB Antigenic variation in relapsing fever species of ***Borrelia*** , including B. hermsii, B. duttonii and B. turicatae, is reviewed. Subjects discussed include virulence properties, clinical and experimental infections, variable antigens, active and silent genes for variable antigens, linear plasmids, and models for the mechanism of ***vmp*** switching.
- TI Antigenic variation in relapsing fever ***Borrelia*** species.
- AB Antigenic variation in relapsing fever species of ***Borrelia*** , including B. hermsii, B. duttonii and B. turicatae, is reviewed. Subjects discussed include virulence properties, clinical and experimental infections, variable antigens, active and silent genes for variable antigens, linear plasmids, and models for the mechanism of ***vmp*** switching.
- BT Spirochaetales; Gracilicutes; bacteria; prokaryotes; ***Borrelia*** ; Spirochaetaceae
- ORGN Spirochaetaceae; ***Borrelia*** hermsii; ***Borrelia*** duttonii; ***Borrelia***
- L4 ANSWER 76 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on $_{\mbox{\scriptsize STN}}$
- AN 1991:104960 BIOSIS <<LOGINID::20090609>>
- DN PREV199140047780; BR40:47780
- TI MULTIPHASIC ANTIGENIC VARIATION IN THE BACTERIUM THAT CAUSES RELAPSING FEVER.
- AU BARBOUR A G [Reprint author]
- CS DEP MICROBIOL, UNIV TEXAS HEALTH SCI CENT, SAN ANTONIO, TEXAS 78284, USA
- SO (1990) pp. 183-200. VAN DER PLOEG, L. H. T., C. R. CANTOR AND H. J. VOGEL (ED.). IMMUNE RECOGNITION AND EVASION: MOLECULAR ASPECTS OF HOST-PARASITE INTERACTION; P AND S BIOMEDICAL SCIENCES SYMPOSIUM, NEW YORK, NEW YORK, USA, JUNE 3-5, 1988. XIII+315P. ACADEMIC PRESS, INC.: SAN DIEGO, CALIFORNIA, USA; LONDON, ENGLAND, UK. ILLUS. ISBN: 0-12-711710-5.
- DT Book Conference; (Meeting)
- FS BR
- LA ENGLISH
- ED Entered STN: 26 Feb 1991 Last Updated on STN: 26 Feb 1991
- IT Miscellaneous Descriptors
 - ***BORRELIA*** HUMAN RAT ***VMP*** GENE OUTER MEMBRANE PROTEIN
- L4 ANSWER 77 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on

STN

- AN 1991:15290 BIOSIS <<LOGINID::20090609>>
- DN PREV199140003620; BR40:3620
- TI ANTIGENIC VARIATION OF A RELAPSING FEVER ***BORRELIA*** SPECIES.
- AU BARBOUR A G [Reprint author]
- CS DEP MICROBIOL MEDICINE, UNIVERSITY TEXAS HEALTH SCIENCE CENTER SAN ANTONIO, SAN ANTONIO, TEXAS 78284, USA
- SO Annu. Rev. Microbiol., (1990) pp. 155-172. ORNSTON, N. L. (ED.). ANNUAL REVIEW OF MICROBIOLOGY, VOL. 44. XIII+748P. ANNUAL REVIEWS INC.: PALO ALTO, CALIFORNIA, USA. ILLUS.

 Publisher: Series: Appual Review of Microbiology

Publisher: Series: Annual Review of Microbiology. CODEN: ARMIAZ. ISSN: 0066-4227. ISBN: 0-8243-1144-2.

- DT Book
- FS BR
- LA ENGLISH
- ED Entered STN: 11 Dec 1990
 Last Updated on STN: 11 Dec 1990
- TI ANTIGENIC VARIATION OF A RELAPSING FEVER ***BORRELIA*** SPECIES.
- IT Miscellaneous Descriptors
 REVIEW OUTER MEMBRANE PROTEIN DNA REARRANGEMENT PLASMID ***VMP***
 GENES
- L4 ANSWER 78 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN DUPLICATE 49
- AN 1990:448185 BIOSIS <<LOGINID::20090609>>
- DN PREV199090098825; BA90:98825
- TI JUXTAPOSITION OF EXPRESSED VARIABLE ANTIGEN GENES WITH A CONSERVED TELOMERE IN THE BACTERIUM ***BORRELIA*** -HERMSII.
- AU KITTEN T [Reprint author]; BARBOUR A G
- CS DEP MICROBIOLOGY, UNIVERSITY TEXAS HEALTH SCI CENTER, SAN ANTONIO, TEXAS 78284, USA
- SO Proceedings of the National Academy of Sciences of the United States of America, (1990) Vol. 87, No. 16, pp. 6077-6081.

 CODEN: PNASA6. ISSN: 0027-8424.
- DT Article
- FS BA
- LA ENGLISH
- ED Entered STN: 7 Oct 1990 Last Updated on STN: 7 Oct 1990
- ***Borrelia*** hermsii, an agent of relapsing fever, survives in AB mammals through antigenic variation. Change in serotype-specific variable outer membrane proteins (***Vmps***) occurs when a ***Vmp*** at an expression site is replaced with a previously silent gene for ***Vmp*** . Silent and active genes are on separate linear plasmids. The upstream site for a nonreciprocal recombination between two linear plasmids is near the 5' ends of the expressed and silent genes. In the present study we sought the downstream recombination sites in two serotypes, 7 and 21. Restriction fragments containing plasmid telomeres were identified by susceptibility to digestion with BAL-31 and rapid reannealment following denaturation. Whereas both silent genes and a minority population of both expression-linked genes were several kilobases from the telomeres, the predominant population of both expressed genes had 3' ends near plasmid telomeres. Sequence analysis of the predominant expression plasmids revealed that the telomeric sequences were the same in serotypes 7 and 21. Identical sequence was also downstream of silent ***Vmp*** genes. Switching of ***Vmp*** genes appears to occur by

recombination that involves both upstream and downstream sites. The

- expression plasmid's telomere is preserved in the recombination event.
- TI JUXTAPOSITION OF EXPRESSED VARIABLE ANTIGEN GENES WITH A CONSERVED TELOMERE IN THE BACTERIUM ***BORRELIA*** -HERMSII.
- ***Borrelia*** hermsii, an agent of relapsing fever, survives in mammals through antigenic variation. Change in serotype-specific variable outer membrane proteins (***Vmps***) occurs when a ***Vmp*** gene at an expression site is replaced with a previously silent gene for another ***Vmp***. Silent and active genes are on separate linear plasmids. The upstream site for a nonreciprocal recombination between two linear plasmids. . . revealed that the telomeric sequences were the same in serotypes 7 and 21. Identical sequence was also downstream of silent ***Vmp*** genes. Switching of ***Vmp*** genes appears to occur by recombination that involves both upstream and downstream sites. The expression plasmid's telomere is preserved in. . .
- L4 ANSWER 79 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN DUPLICATE 50
- AN 1991:31053 BIOSIS <<LOGINID::20090609>>
- DN PREV199191020404; BA91:20404
- TI THE VARIABLE ANTIGENS ***VMP7*** AND ***VMP21*** OF THE RELAPSING FEVER BACTERIUM ***BORRELIA*** -HERMSII ARE STRUCTURALLY ANALOGOUS TO THE VSG PROTEINS OF THE AFRICAN TRYPANOSOME.
- AU BURMAN N [Reprint author]; BERGSTROM S; RESTREPO B I; BARBOUR A G
- CS DEP MICROBIOL, UNIV UMEA, S-901 87 UMEA, SWEDEN
- SO Molecular Microbiology, (1990) Vol. 4, No. 10, pp. 1715-1726. CODEN: MOMIEE. ISSN: 0950-382X.
- DT Article
- FS BA
- LA ENGLISH
- OS GENBANK-X53926; GENBANK-X53927
- ED Entered STN: 3 Jan 1991 Last Updated on STN: 3 Jan 1991
- The relapsing fever agent ***Borrelia*** hermsii avoids the host's AΒ immune response by the strategy of multiphasic antigenic variation. A given ***Borrelia*** cell can express one of a number of alleles for polymorphic outer-membrane proteins, known as ***Vmp*** proteins. The genes for the variant-specific ***Vmp*** proteins of serotypes 7 and genes for the variant-specific variant strain HS1 were sequenced. The genes, which were designated ***vmp7*** and ***vmp21*** , were obtained from populations of ***borreliae*** before and after a switch in serotypes from 7 to 21. The analysis showed that ***vmp7*** and are 77% identical in terms of their coding sequence. The deduced translation products of ***vmp7*** and ***vmp21*** polypeptides of 369 (37.2kD) and 364 amino acids (37.1kD), respectively. ***Vmp7*** and ***Vmp21*** have sequence features of prokaryotic lipoproteins and are processed as such during expression in E. coli. The secondary structure predictions of the ***Vmp*** proteins reveals analogous structures to the VSG proteins of the African trypanosome.
- TI THE VARIABLE ANTIGENS ***VMP7*** AND ***VMP21*** OF THE RELAPSING FEVER BACTERIUM ***BORRELIA*** -HERMSII ARE STRUCTURALLY ANALOGOUS TO THE VSG PROTEINS OF THE AFRICAN TRYPANOSOME.
- AB The relapsing fever agent ***Borrelia*** hermsii avoids the host's immune response by the strategy of multiphasic antigenic variation. A given ***Borrelia*** cell can express one of a number of alleles for polymorphic outer-membrane proteins, known as ***Vmp*** proteins. The genes for the variant-specific ***Vmp*** proteins of serotypes 7 and 21 of B. hermsii strain HS1 were sequenced. The genes, which were

designated ***vmp7*** and ***vmp21*** , were obtained from populations of ***borreliae*** before and after a switch in serotypes from 7 to 21. The analysis showed that ***vmp7*** and ***vmp21*** are 77% identical in terms of their coding sequence. The deduced translation products of ***vmp7*** and ***vmp21*** polypeptides of 369 (37.2kD) and 364 amino acids (37.1kD), respectively. ***Vmp7*** and ***Vmp21*** have sequence features of prokaryotic lipoproteins and are processed as such during expression in E. coli. The secondary structure predictions of the ***Vmp*** proteins reveals analogous structures to the VSG proteins of the African trypanosome. ANSWER 80 OF 87 CAPLUS COPYRIGHT 2009 ACS on STN 1991:605225 CAPLUS <<LOGINID::20090609>> 115:205225 OREF 115:35001a,35004a Multiphasic antigenic variation in the bacterium that causes relapsing fever Barbour, Alan G. Health Sci. Cent., Univ. Texas, San Antonio, TX, 78284, USA Immune Recognit. Evasion: Mol. Aspects Host-Parasite Interact. (1990), 183-99. Editor(s): Van der Ploeg, Lex H. T.; Cantor, Charles R.; Vogel, Henry J. Publisher: Academic, San Diego, Calif. CODEN: 57AEAO Conference; General Review English A review with 27 refs. of multiphasic antigenic variation in ***Borrelia*** and the role of recombination in ***vmp*** (variable major protein) gene switching. A review with 27 refs. of multiphasic antigenic variation in ***Borrelia*** and the role of recombination in ***vmp*** (variable major protein) gene switching. review antigen variation ***Borrelia*** ; gene switching antigen ***Borrelia*** review ***Borrelia*** (antigenic variation in) Gene and Genetic element, microbial RL: BIOL (Biological study) (for variable antigens, switching of, in antigenic variation in ***Borrelia***) Recombination, genetic (in antigenic variation, in ***Borrelia***) Antigens RL: BIOL (Biological study) (variability of, in ***Borrelia***) ANSWER 81 OF 87 CABA COPYRIGHT 2009 CABI on STN DUPLICATE 51 19920512260 Antigenic variation of a relapsing fever ***Borrelia*** species Barbour, A. G. Department of Microbiology and Medicine, University of Texas Health Science Center at San Antonio, San Antonio, TX 78284, USA. Annual Review of Microbiology, (1990) Vol. 44, pp. 155-171. 39 ref. Publisher: Annual Reviews Inc. Palo Alto, California ISSN: 0066-4227; ISBN: 0-8243-1144-2 United States

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Journal

- LA English
- ED Entered STN: 1 Nov 1994
 Last Updated on STN: 1 Nov 1994
- AB This review examines the immunology of relapsing fever and antigenic variation in B. hermsii. The variable antigens of this spirochaete are outer membrane proteins, and antigenic variation is the consequence of DNA rearrangements. The ***vmp*** genes (of Variable Major Proteins) are located on linear plasmids. The activation of a new ***vmp*** is the result of recombination between different linear plasmids.
- TI Antigenic variation of a relapsing fever ***Borrelia*** species.
- AB . . . The variable antigens of this spirochaete are outer membrane proteins, and antigenic variation is the consequence of DNA rearrangements. The ***vmp*** genes (of Variable Major Proteins) are located on linear plasmids. The activation of a new ***vmp*** is the result of recombination between different linear plasmids.
- BT Spirochaetales; Gracilicutes; bacteria; prokaryotes; ***Borrelia***; Spirochaetaceae
- ORGN Spirochaetaceae; ***Borrelia*** hermsii
- L4 ANSWER 82 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN
- AN 1989:299884 BIOSIS <<LOGINID::20090609>>
- DN PREV198937014261; BR37:14261
- TI ANTIGENIC VARIATION IN RELAPSING FEVER ***BORRELIA*** SPECIES GENETICS ASPECTS.
- AU BARBOUR A [Reprint author]
- CS DEP OF MICROBIOL, UNIV OF TEX HEALTH SCI CENT, 7703 FLOYD CURL DRIVE, SAN ANTONIO, TEX 78284-7758, USA
- SO (1989) pp. 783-790. BERG, D. E. AND M. M. HOWE (ED.). MOBILE DNA. XVII+972P. AMERICAN SOCIETY FOR MICROBIOLOGY: WASHINGTON, D.C., USA. ILLUS. MAPS. ISBN: 1-55581-005-5.
- DT Book
- FS BR
- LA ENGLISH
- ED Entered STN: 27 Jun 1989 Last Updated on STN: 27 Jun 1989
- TI ANTIGENIC VARIATION IN RELAPSING FEVER ***BORRELIA*** SPECIES GENETICS ASPECTS.
- IT Miscellaneous Descriptors
 REVIEW OUTER MEMBRANE PROTEINS DNA REARRANGEMENT ***VMP*** GENES
 SWITCHING
- L4 ANSWER 83 OF 87 LIFESCI COPYRIGHT 2009 CSA on STN
- AN 89:7791 LIFESCI <<LOGINID::20090609>>
- TI Antigenic variation in relapsing fever ***Borrelia*** species:
 Genetic aspects.
 MOBILE DNA.
- AU Barbour, A.; Berg, D.E. [editor]; Howe, M.M. [editor]
- CS Dep. Microbiol., Univ. Texas Health Sci. Cent., 7703 Floyd Curl Dr., San Antonio, TX 78284-7758, USA
- SO (1989) pp. 783-790. ISBN: 1-55581-005-5.
- DT Book
- TC General Review
- FS J; G
- LA English

- AB The following aspects are covered in this review: biology and immunology of relapsing fever ***borreliae***; outer membrane proteins are determinants of serotype specificity; antigenic variation is the consequence of DNA rearrangements; the ***vmp*** genes are on linear plasmids; and possible mechanisms of ***vmp*** gene switching.
- TI Antigenic variation in relapsing fever ***Borrelia*** species:

 Genetic aspects.

 MOBILE DNA.
- AB The following aspects are covered in this review: biology and immunology of relapsing fever ***borreliae***; outer membrane proteins are determinants of serotype specificity; antigenic variation is the consequence of DNA rearrangements; the ***vmp*** genes are on linear plasmids; and possible mechanisms of ***vmp*** gene switching.
- UT reviews; membrane proteins; genes; ***Borrelia*** ; antigenic variants;
 genetic variance; ***vmp*** gene
- L4 ANSWER 84 OF 87 EMBASE COPYRIGHT (c) 2009 Elsevier B.V. All rights reserved on STN
- AN 1988223362 EMBASE <<LOGINID::20090609>>
- TI Genetic mechanisms of bacterial antigenic variation.
- AU Seifert, H.S.; So, M.
- CS Department of Microbiology and Immunology, Northwestern Medical and Dental Schools, Chicago, IL 60611, United States.
- SO Microbiological Reviews, (1988) Vol. 52, No. 3, pp. 327-336. ISSN: 0146-0749 CODEN: MBRED3
- CY United States
- DT Journal; General Review; (Review)
- FS 026 Immunology, Serology and Transplantation 004 Microbiology: Bacteriology, Mycology, Parasitology and Virology
- LA English
- SL English
- ED Entered STN: 11 Dec 1991 Last Updated on STN: 11 Dec 1991
- AΒ The studies described above indicate that procaryotes have evolved a variety of mechanisms to vary their surface coats. N. gonorrhoeae primarily uses DNA transformation to effect pilus antigenic variation at the recombinational level. It also uses recombination (and perhaps also DNA transformation) to bring about P.II antigenic variation at the translational level. Finally, ***Borrelia*** organisms have evolved a plasmid recombination system to undergo ***VMP*** antigenic variation. To place procaryotic antigenic variation into proper perspective, we end this review with a brief consideration of the host immune system. Mammals have also evolved what could be considered an antigenic variation system, i.e., the generation of antibodies with different antigen-binding specificities. The arrangement of multiple copies of V, D, and J gene segments in the mammalian genome is reminiscent of the arrangement of silent pilin gene segments in the gonococcal chromosome. However, unlike ***VMP*** expression, the generation of a functional pilin, P.II, and expressing immunoglobulin gene does not involve expression sites. Instead, a complete immunoglobulin gene is created by recombinational joining of various gene segments, with concomitant deletion of intervening sequences. A system that appears to resemble the gonococcal pilin mechanism has been described for chicken immunoglobulin light chains. The light chain variants all are derived from a unique V-J rearrangement, with diversification occurring by gene conversion from other V gene copies to this single expressed gene within the Bursa of Fabricius. Four main processes appear to be responsible for the generation of antibody

diversity in mammalian cells. The first, known as 'combinational diversity', is the joining of V and J gene segments in various combinations. Diversity could also be generated by imprecise joining at V-J, V-D, and D-J junctions. In addition, joining of the V(H)-D and D-J(H) segments could lead to insertion of one to several nucleotides at these junctions. Finally, sequence changes could occur in immunoglobulin gene segments by somatic mutation. Whether these four processes also contribute to antigenic variation in procaryotic systems is not known at present. Since both the procaryotic and eucaryotic systems operate at the recombinational level, it is possible that the first three processes which contribute to immunoglobulin diversity also play a role in procaryotic antigenic variation. As for somatic mutations, it is clear that antigenic drift contributes significantly to the generation of hemagglutinin and neuraminidase variants of the flu virus. It is therefore likely that this process also contributes to sequence variability of the pilin, P.II, and ***VMP*** genes. In addition, gene conversion is thought to contribute to the generation of somatic mutation in immunoglobulin genes. In summary, it is interesting to note that the systems of antigenic variation and immunoglobulin diversification have evolved in a similar and complementary fashion, with DNA recombination playing a central mechanistic role. It is highly likely that the two systems developed together, with each providing the evolutionary pressure needed by the other. Finally, the examples of antigenic variation covered in this review illustrate the fascinating and diverse ways microbes have found to regulate and alter gene expression.

. . It also uses recombination (and perhaps also DNA transformation) AB to bring about P.II antigenic variation at the translational level. ***Borrelia*** organisms have evolved a plasmid recombination Finally, ***VMP*** antigenic variation. To place procaryotic system to undergo antigenic variation into proper perspective, we end this review with a brief consideration of the. . . genome is reminiscent of the arrangement of silent pilin gene segments in the gonococcal chromosome. However, unlike pilin, P.II, and ***VMP*** expression, the generation of a functional expressing immunoglobulin gene does not involve expression sites. Instead, a complete immunoglobulin gene is. . . the flu virus. It is therefore likely that this process also contributes to sequence variability of the pilin, P.II, and ***VMP*** genes. In addition, gene conversion is thought to contribute to the generation of somatic mutation in immunoglobulin genes. In summary,.

- L4 ANSWER 85 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN DUPLICATE 52
- AN 1985:383147 BIOSIS <<LOGINID::20090609>>
- DN PREV198580053139; BA80:53139
- TI VARIABLE MAJOR PROTEINS OF ***BORRELIA*** -HERMSII EPITOPE MAPPING AND PARTIAL SEQUENCE ANALYSIS OF CYANOGEN BROMIDE PEPTIDES.
- AU BARSTAD P A [Reprint author]; COLIGAN J E; RAUM M G; BARBOUR A G
- CS LAB MICROBIAL STRUCTURE FUNCTION, ROCKY MOUNTAIN LAB, NATL INST ALLERGY INFECTIOUS DISEASES, HAMILTON, MONTANA 59840, USA
- SO Journal of Experimental Medicine, (1985) Vol. 161, No. 6, pp. 1302-1314. CODEN: JEMEAV. ISSN: 0022-1007.
- DT Article
- FS BA
- LA ENGLISH
- AB The variable major proteins (***VMP***) of serotypes 7 and 21 of the relapsing fever agent B. hermsii were isolated by detergent extraction and high performance liquid chromatography. CNBr digestion of the isolated

yielded 2 peptides of apparent MW 20,000 (20 K) and 16 K from ***VMP7*** , and 3 peptides of 14.5, 14, and 7 K MW from ***VMP21*** Serotype-specific monoclonal antibodies bound in Western blots to 1 of ***VMP*** . A each of the 2 or 3 CNBr fragments from the homologous single monoclonal antibody bound to the whole cells, the isolated ***VMP*** , and a CNBr fragment of both serotype 7 and serotype 21. (This cross-reactive antibody did not, however, bind to any of 4 other serotypes examined). Regional conservation of structure between ***VMP7*** and ***VMP21*** was also shown by amino acid sequence analysis of the N-termini of the 5 CNBr fragments. One pair of aligned fragments from ***VMP7*** and ***VMP21*** had 80% amino acid homologies between 2 ***VMP*** suggest that these proteins are products of members of a polygene family. VARIABLE MAJOR PROTEINS OF ***BORRELIA*** -HERMSII EPITOPE MAPPING AND ТΤ PARTIAL SEQUENCE ANALYSIS OF CYANOGEN BROMIDE PEPTIDES. The variable major proteins (***VMP****) of serotypes 7 and 21 of the AB relapsing fever agent B. hermsii were isolated by detergent extraction and high performance liquid chromatography. CNBr digestion of the isolated ***VMP*** yielded 2 peptides of apparent MW 20,000 (20 K) and 16 K from Serotype-specific monoclonal antibodies bound in Western blots to 1 of each of the 2 or 3 CNBr fragments from the homologous ***VMP*** . A single monoclonal antibody bound to the whole cells, the isolated ***VMP*** , and a CNBr fragment of both serotype 7 and serotype 21. (This cross-reactive antibody did not, however, bind to any of 4 other serotypes examined). Regional conservation of structure between ***VMP7*** and ***VMP21*** was also shown by amino acid sequence analysis of the ${\tt N-termini}$ of the 5 CNBr fragments. One pair of aligned fragments from ***VMP7*** and ***VMP21*** had 80% amino acid homologies between 2***VMP*** suggest that these proteins are products of members of a polygene family. ANSWER 86 OF 87 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on L4DUPLICATE 53 1986:142045 BIOSIS <<LOGINID::20090609>> ΑN PREV198681052461; BA81:52461 DN TRANSPOSITION OF STRUCTURAL GENES TO AN EXPRESSION SEQUENCE ON A LINEAR PLASMID CAUSES ANTIGENIC VARIATION IN THE BACTERIUM ***BORRELTA*** -HERMSII. ΑU PLASTERK R H A [Reprint author]; SIMON M I; BARBOUR A G CS CALIFORNIA INST TECHNOLOGY, DIV BIOLOGY 147-75 PASADENA, CALIFORNIA 91125, SO Nature (London), (1985) Vol. 318, No. 6043, pp. 257-263. CODEN: NATUAS. ISSN: 0028-0836. DT Article FS LA ENGLISH ED Entered STN: 25 Apr 1986 Last Updated on STN: 25 Apr 1986 AΒ In ***Borrelia*** hermsii, a spirochaete that causes relapsing fever, the switch between expression of two frequent variable major protein (***VMP***) types (7 and 21) is associated with a DNA rearrangement. Both cell types 7 and 21 contain untranscribed 7 and 21 ***VMP***

- genes on linear plasmids. The serotype 7 cells contain an additional copy of the 7 ***VMP*** gene fused to an expression sequence on another linear plasmid. Switching to the 21 serotype involves removal of the transcribed 7 ***VMP*** gene and fusion of a copy of the 21 ***VMP*** gene to this same expression sequence. Thus, recombination between linear plasmids can activate different ***VMP*** genes.
- TI TRANSPOSITION OF STRUCTURAL GENES TO AN EXPRESSION SEQUENCE ON A LINEAR PLASMID CAUSES ANTIGENIC VARIATION IN THE BACTERIUM ***BORRELIA***
 -HERMSII.
- L4 ANSWER 87 OF 87 EMBASE COPYRIGHT (c) 2009 Elsevier B.V. All rights reserved on STN
- AN 1986060790 EMBASE <<LOGINID::20090609>>
- TI Transposition of structural genes to an expression sequence on a linear plasmid causes antigenic variation in the bacterium ***Borrelia*** hermsii.
- AU Plasterk, R.H.A.; Simon, M.I.; Barbour, A.G.
- CS California Institute of Technology, Division of Biology 147-75, Pasadena, CA 91125, United States.
- SO Nature, (1985) Vol. 31 B, No. 6043, pp. 257-263. ISSN: 0028-0836 CODEN: NATUAS
- CY United Kingdom
- DT Journal
- FS 022 Human Genetics
 - O26 Immunology, Serology and Transplantation
 - 004 Microbiology: Bacteriology, Mycology, Parasitology and Virology
- LA English
- ED Entered STN: 10 Dec 1991 Last Updated on STN: 10 Dec 1991
- TI Transposition of structural genes to an expression sequence on a linear plasmid causes antigenic variation in the bacterium ***Borrelia*** hermsii.

of the 7 $\ \ ^{***VMP***}$ gene fused to an expression sequence on another linear plasmid. Switching to the 21 serotype involves removal of the transcribed 7 ***VMP*** gene and fusion of a copy of the 21 ***VMP*** gene to this same expression sequence. Thus recombination between linear plasmids can activate different ***VMP*** genes. CT Medical Descriptors: ****borrelia hermsii*** gene translocation heredity nonhuman *plasmid priority journal *serotype *antigen *bacterial antigen

An alternative to view hit terms when display exceeds KWIC processing limits is to use HIT display format.